-key terms

10/034500

FILE 'HCAPLUS' ENTERED AT 11:20:12 ON 10 APR 2003 L1 6 S (LAWSON? OR L) (W) INTRACELL? AND (OMP OR OUTER MEMBRAN? PROTEIN) 14 S (LAWSON? OR L) (W) INTRACELL? AND ANTIGEN? L215 S L1 OR L2 L3 ANSWER 1 OF 15 HCAPLUS COPYRIGHT 2003 ACS L3 2002:503432 HCAPLUS ACCESSION NUMBER: DOCUMENT NUMBER: 137:77871 Cloning of genes for novel Lawsonia TITLE: intracellularis outer membrane proteins and their use in preparing vaccines for porcine proliferative enteropathy Jacobs, Antonius A. C.; Vermeij, Paul INVENTOR(S): Akzo Nobel N.V., Neth. PATENT ASSIGNEE(S): Eur. Pat. Appl., 26 pp. SOURCE: CODEN: EPXXDW DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE _____ ______ ---------20020703 EP 2001-204919 20011214 EP 1219711 A2 20021106 А3 EP 1219711 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR JP 2003000276 A2 20030107 JP 2001-385373 20011219 20011220 20020627 AU 2001-97371 AU 2001097371 Α5 EP 2000-204660 A 20001220 PRIORITY APPLN. INFO.: The present invention relates i.a. to nucleic acid sequences encoding novel Lawsonia intracellularis proteins. It furthermore relates to DNA fragments, recombinant DNA mols. and live recombinant carriers comprising these sequences. Also it relates to host cells comprising such nucleic acid sequences, DNA fragments, recombinant DNA mols. and live recombinant carriers. Moreover, the invention relates to proteins encoded by these nucleotide sequences. The invention also relates to vaccines for combating Lawsonia intracellularis infections and methods for the prepn. thereof. Finally the invention relates to diagnostic tests for the detection of Lawsonia intracellularis DNA, the detection of Lawsonia intracellularis antigens and of antibodies against Lawsonia intracellularis. ANSWER 2 OF 15 HCAPLUS COPYRIGHT 2003 ACS 2002:415165 HCAPLUS ACCESSION NUMBER: 137:137337 DOCUMENT NUMBER: LsaA, an antigen involved in cell TITLE: attachment and invasion, is expressed by Lawsonia intracellularis during infection in vitro and in vivo McCluskey, Jackie; Hannigan, Joanne; Harris, AUTHOR(S): Jennifer D.; Wren, Brendan; Smith, David G. E.

Searcher: Shears 308-4994

CORPORATE SOURCE:

Zoonotic & Animal Pathogens Research Laboratory,

Department of Medical Microbiology, University

of Edinburgh, Edinburgh, UK

SOURCE: Infection and Immunity (2002), 70(6), 2899-2907

CODEN: INFIBR; ISSN: 0019-9567

PUBLISHER: American Society for Microbiology

DOCUMENT TYPE: Journal LANGUAGE: English

AB Lawsonia intracellularis has been identified

recently as the etiol. agent of proliferative enteropathies, which are characterized by intestinal epithelial hyperplasia and assocd. moderate immune responses. This disease complex has been reported in a broad range of animals prevalently in pigs. and I

in a broad range of animals, prevalently in pigs, and L. intracellularis has been linked with ulcerative colitis in humans. L. intracellularis is an obligate

intracellular bacterium, and the pathogenic mechanisms used to cause disease are unknown. Using in vitro-grown organisms as a source of genomic DNA, we identified a Lawsonia gene which encodes a surface antigen, LsaA (for Lawsonia surface antigen),

assocd. with attachment to and entry into cells. The deduced amino acid sequence of this protein showed some similarity to members of a novel protein family identified in a no. of other bacterial

pathogens but for which roles are not fully defined. Transcription of this gene was detected by reverse transcription-PCR in ${\bf L}$

. intracellularis grown in vitro in IEC18 cells and in bacteria present in ileal tissue from infected animals. Immunohistochem. with specific monoclonal antibody and

immunoblotting with sera from infected animals demonstrated that

LsaA protein is synthesized by L. intracellularis

during infection. Expression of this gene during infection in vitro and in vivo suggests that this surface **antigen** is involved

during infection, and phenotypic anal. indicated a role during L. intracellularis attachment to and entry into

intestinal epithelial cells.

REFERENCE COUNT: 48 THERE ARE 48 CITED REFERENCES AVAILABLE

FOR THIS RECORD. ALL CITATIONS AVAILABLE

IN THE RE FORMAT

L3 ANSWER 3 OF 15 HCAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2002:368499 HCAPLUS

DOCUMENT NUMBER: 136:382847

TITLE: Genes for antigenic proteins of

Lawsonia and their use diagnosis and prophylaxis

of Lawsonia infection

INVENTOR(S): Rosey, Everett Lee; King, Kendall Wayne; Good,

Robert Trygve; Strugnell, Richard Anthony

PATENT ASSIGNEE(S): Agriculture Victoria Services Pty. Ltd.,

Australia; Australian Pork Limited; Pfizer

Products, Inc.

SOURCE: PCT Int. Appl., 155 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE: En FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2002038594 A1 20020516 WO 2001-AU1462 20011109
WO 2002038594 C2 20021107

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W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH,
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             NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ,
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             TD, TG
     AU 2002014810
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                                          AU 2002-14810
                                                           20011109
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PRIORITY APPLN. INFO.:
                                       AU 2000-1381
                                                    A 20001110
                                       US 2000-249596P P 20001117
                                       WO 2001-AU1462 W 20011109
     The present invention relates generally to therapeutic compns. for
AB
     the treatment and/or prophylaxis of intestinal disease conditions in
     animals and birds caused or exacerbated by Lawsonia
     intracellularis or similar or otherwise related
     microorganisms. In particular, the present invention provides a
     novel gene derived from Lawsonia intracellularis
     , which encodes an immunogenic polypeptide that is particularly
     useful as an antigen in a vaccine prepn. for conferring
     humoral immunity against Lawsonia intracellularis
     and related pathogens in animal hosts, wherein said polypeptide is
     selected from the group consisting of flhB, fliR, ntrC, glnH, motA,
    motB, tlyC, ytfM, and ytfN polypeptides, or a homolog, analog or
     deriv. of any one or more of said polypeptides. The present
     invention is also directed to methods for the treatment and/or
    prophylaxis of such intestinal disease conditions and to diagnostic
     agents and procedures for detecting Lawsonia
     intracellularis or similar or otherwise related
    microorganisms.
REFERENCE COUNT:
                         3
                              THERE ARE 3 CITED REFERENCES AVAILABLE FOR
                              THIS RECORD. ALL CITATIONS AVAILABLE IN
                              THE RE FORMAT
    ANSWER 4 OF 15 HCAPLUS COPYRIGHT 2003 ACS
                        2002:256061 HCAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                        136:261820
                        Swine vaccines for proliferative ileitis
TITLE:
                        comprising Lawsonia
                        intracellularis antigens
                        Arizona Board of Regents on Behalf of the
PATENT ASSIGNEE(S):
                        University of Arizona, USA
SOURCE:
                        PCT Int. Appl., 43 pp.
                        CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                        English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                                          APPLICATION NO. DATE
     PATENT NO.
                     KIND DATE
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                    A2 20020404 WO 2001-US30284 20010927
     WO 2002026250
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             HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
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LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT,
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             TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
                                             AU 2001-93151
     AU 2001093151
                        A5
                             20020408
                                                               20010927
                                          US 2000-677108
PRIORITY APPLN. INFO.:
                                                            Α
                                                               20000929
                                          WO 2001-US30284 W 20010927
     A proliferative ileitis vaccine comprising tissue culture grown
AΒ
     Lawsonia intracellularis and methods of making
     said vaccines. Proliferative ileitis vaccines described include
     those contg. whole L. intracellularis, exts. of
     L. intracellularis, protective immunogenic submits
     of L. intracellularis, recombinant immunogens of
     L. intracellularis and naked DNA of L.
     intracellularis. The vaccines of this invention may be
     inactivated or modified live and contain adjuvants and/or
     stabilizers. The vaccines of this invention may be in a liq. or
     lyophilized form. Also disclosed are monoclonal antibodies which
     neutralize the growth of L. intracellularis and
     which may be used for diagnosing proliferative ileitis as well as
     for quantitating antigen during vaccine prodn.
     ANSWER 5 OF 15 HCAPLUS COPYRIGHT 2003 ACS
L3
                          2001:297553 HCAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                          134:321599
TITLE:
                          Cloning of Lawsonia genes htrA, ponA, hypC,
                          lysS, ycfW, abcl, and omp100, their encoded
                          proteins or peptides and therapeutic use in
                          diagnosis and as vaccine
INVENTOR(S):
                          Rosey, Everett Lee
PATENT ASSIGNEE(S):
                          Pfizer Products Inc., USA
SOURCE:
                          Eur. Pat. Appl., 80 pp.
                          CODEN: EPXXDW
DOCUMENT TYPE:
                          Patent
                          English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                       KIND DATE
                                             APPLICATION NO.
                                                               DATE
     PATENT NO.
                       ____
     EP 1094070
                       A2
                             20010425
                                             EP 2000-309125
                                                               20001017
                             20020109
     EP 1094070
                       A3
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,
             PT, IE, SI, LT, LV, FI, RO
                                             JP 2000-320736
                                                               20001020
     JP 2001169787
                       A2
                             20010626
                                             US 2002-210296
     US 2003021802
                        Α1
                             20030130
                                                               20020801
PRIORITY APPLN. INFO.:
                                          US 1999-160922P P
                                                               19991022
                                          US 1999-163858P
                                                            P
                                                               19991105
                                          US 2000-689065
                                                           A1 20001012
     The present invention relates generally to therapeutic compns. for
AB
     the treatment and/or prophylaxis of intestinal disease conditions in
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The present invention relates generally to therapeutic compns. For the treatment and/or prophylaxis of intestinal disease conditions in pigs or other animals caused or exacerbated by Lawsonia intracellularis or similar or otherwise related microorganism, such as porcine proliferative enteropathy (PPE). In particular, the present invention provides novel genes htrA, ponA,

hypC, lysS, ycfW, abcl, and omp100 derived from Lawsonia intracellularis genomic regions A and B. These genes encode sequence homologs to lysyl-tRNA synthetase (gene lysS), transmembrane or integral membrane protein (abcl), hydrogenase maturation protein (hypC), penicillin binding protein (ponA), and periplasmic serine protease protein (htrA) resp. The invention also relates to constructing these gene expression vector to produce recombinant protein using E. coli. Methods of expressing recombinant htrA and omp100 proteins in E. coli are also provided. The invention also provides the immunogenic peptides or proteins encoded by these genes that are particularly useful as an antigen in vaccine prepn. for conferring humoral immunity against Lawsonia intracellularis and related pathogens in animal hosts. The present invention is also directed to methods for the treatment and/or prophylaxis of such intestinal disease conditions and to diagnostic agents and procedures for detecting Lawsonia intracellularis or similar or otherwise related microorganisms.

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ANSWER 6 OF 15 HCAPLUS COPYRIGHT 2003 ACS
                       2000:824297 HCAPLUS
ACCESSION NUMBER:
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DOCUMENT NUMBER: 134:1364

Lawsonia-derived gene tlyA and related hemolysin TITLE:

polypeptides, peptides and proteins and their uses for diagnosis and treatment of avian and

porcine infections

Panaccio, Michael; Rosey, Everett Lee; Hasse, INVENTOR(S):

Detlef; Ankenbauer, Robert Gerard

Pfizer Products Inc, USA; Agriculture Victoria PATENT ASSIGNEE(S):

Services Pty Ltd; Pig Research and Development

Corporation

SOURCE: PCT Int. Appl., 86 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PATENT NO.
                      KIND DATE
                                                  APPLICATION NO. DATE
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     WO 2000069906
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               BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                                EP 2000-924978
     EP 1177213
                         A1 20020206
                                                                      20000511
          R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,
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PRIORITY APPLN. INFO.:
                                               US 1999-134022P P 19990513
                                                                W 20000511
                                               WO 2000-AU439
     The present invention relates generally to therapeutic compns. for
     the treatment and/or prophylaxis of intestinal disease conditions in
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Shears 308-4994 Searcher :

animals and birds caused or exacerbated by Lawsonia

intracellularis or similar or otherwise related
microorganism. In particular, the present invention provides a
novel gene derived from Lawsonia intracellularis
which encodes an immunogenic TylA hemolysin peptide, polypeptide or
protein that is particularly useful as an antigen in
vaccine prepn. for conferring humoral immunity against
Lawsonia intracellularis and related pathogens in
animal hosts. The present invention is also directed to methods for
the treatment and/or prophylaxis of such intestinal disease
conditions and to diagnostic agents and procedures for detecting
Lawsonia intracellularis or similar or otherwise
related microorganisms.

REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 7 OF 15 HCAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2000:824296 HCAPLUS

DOCUMENT NUMBER: 134:14022

TITLE: Lawsonia-derived gene ompH and related

outer membrane protein

H polypeptides, peptides and proteins and their uses for diagnosis and treatment of avian and

porcine infections

INVENTOR(S): Hasse, Detlef; Panaccio, Michael; Sinistaj, Meri

PATENT ASSIGNEE(S): Pig Research and Development Corporation,

Australia; Agriculture Victoria Services Pty Ltd

PCT Int. Appl., 85 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

SOURCE:

| | PAT | ENT I | NO. | | KI | ND ! | DATE | | | A | PPLI | CATI | ои ис | ο. | DATE | | |
|------|------|-------|------|------|--------|------|-------|------|------|------|------|------|----------|-----|-------|-------|-----|
| | WO | 20000 | 0699 | 05 | A: | 1 : | 2000: | 1123 | | W | 0 20 | 00-A | U438 | | 20000 |)511 | |
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| | | | LS, | LT, | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | NO, | NZ, | PL, | PT, |
| | | | RO, | RU, | SD, | SE, | SG, | SI, | SK, | SL, | ТJ, | TM, | TR, | TT, | ΤZ, | UA, | ŪĠ, |
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| | ΕP | 1183 | 268 | | A | 1 : | 20020 | 0306 | | E | P 20 | 00-9 | 2497 | 7 | 20000 |)511 | |
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| | BR | 2000 | 0112 | 90 | A | | 2002 | 0521 | | B | R 20 | 00-1 | 1290 | | 20000 | 0511 | |
| PRIO | RITY | APP | LN. | INFO | . : | | | | | US 1 | 999- | 1339 | 86P | P | 19990 | 0513 | |
| | | | | | | | | | | WO 2 | 000- | AU43 | 8 | W | 2000 |)511 | |
| AB | The | pre | sent | inv | enti | on r | elate | es a | ener | allv | to | ther | apeut | tic | compi | ns. : | for |

AB The present invention relates generally to therapeutic compns. for the treatment and/or prophylaxis of intestinal disease conditions in animals and birds caused or exacerbated by Lawsonia intracellularis or similar or otherwise related microorganism. In particular, the present invention provides a novel gene derived from Lawsonia intracellularis

which encodes an immunogenic OmpH outer membrane peptide, polypeptide or protein that is particularly useful as an antigen in vaccine prepn. for conferring humoral immunity against Lawsonia intracellularis and related pathogens in animal hosts. The present invention is also directed to methods for the treatment and/or prophylaxis of such intestinal disease conditions and to diagnostic agents and procedures for detecting Lawsonia intracellularis or similar or otherwise related microorganisms.

REFERENCE COUNT: THERE ARE 3 CITED REFERENCES AVAILABLE FOR 3 THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 8 OF 15 HCAPLUS COPYRIGHT 2003 ACS L3 2000:824295 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 133:359825

TITLE: Lawsonia-derived gene flgE and related flagellar

hook polypeptides, peptides and proteins and their uses for diagnosis and treatment of avian

and porcine infections

Panaccio, Michael; Rosey, Everett Lee; Sinistaj, INVENTOR(S):

Meri; Hasse, Detlef; Parsons, Jim; Ankenbauer,

Robert Gerard

PATENT ASSIGNEE(S): Pfizer Products Inc., USA; Agriculture Victoria

Services Pty Ltd; Pig Research and Development

Corporation

SOURCE: PCT Int. Appl., 97 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATI | ENT 1 | 10. | | KI | D | DATE | | | I | APPLI | CATI | N NC | ο. | DATE | | |
|----------|-------|------|------|--------|-----|-------|------|-----|--------|-------|-----------|------|-----|------|------|-----|
| WO 2 | 20000 | 0699 | 04 | A: | 1 . | 2000: | 1123 | | - V | 10 20 | IA-00 | u437 | | 2000 | 0511 | |
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| | | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EE, | ES, | FI, | GB, | GD, | GE, | GH, | GM, |
| | | HR, | HU, | ID, | IL, | IN, | IS, | JP, | ΚE, | KG, | ΚP, | KR, | ΚZ, | LC, | LK, | LR, |
| | | LS, | LT, | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | NO, | NZ, | PL, | PT, |
| | | RO, | RU, | SD, | SE, | SG, | SI, | SK, | SL, | ТJ, | TM, | TR, | TT, | TZ, | UA, | UG, |
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| | | ВJ, | CF, | CG, | CI, | CM, | GA, | GN, | GW, | ML, | MR, | ΝE, | SN, | TD, | TG | |
| BR 2 | 20000 | 112 | 94 | A | | 20020 | 0226 | | E | R 20 | 00-1 | 1294 | | 2000 | 0511 | |
| EP : | 11813 | 315 | | A: | 1 . | 20020 | 0227 | | E | P 20 | 00-9 | 2497 | 6 | 2000 | 0511 | |
| | R: | AT, | BE, | CH, | DĒ, | DK, | ES, | FR, | GB, | GR, | IT, | LI, | LU, | NL, | SE, | MC, |
| | | PT, | IE, | SI, | LT, | LV, | FI, | RO | | | | | | | | |
| PRIORITY | APPI | IN. | INFO | . : | | | | | US 1 | .999- | 1339 | 73P | P | 1999 | 0513 | |
| | | | | | | | | | WO 2 | 2000- | AU43 | 7 | W | 2000 | 0511 | |

The present invention relates generally to therapeutic compns. for AB the treatment and/or prophylaxis of intestinal disease conditions in animals and birds caused or exacerbated by Lawsonia intracellularis or similar or otherwise related microorganism. In particular, the present invention provides a novel gene derived from Lawsonia intracellularis which encodes an immunogenic FlgE flagellar hook peptide,

> Shears 308-4994 Searcher :

polypeptide or protein that is particularly useful as an antigen in vaccine prepn. for conferring humoral immunity against Lawsonia intracellularis and related pathogens in animal hosts. The present invention is also directed to methods for the treatment and/or prophylaxis of such intestinal disease conditions and to diagnostic agents and procedures for detecting Lawsonia intracellularis or similar or otherwise related microorganisms.

REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 9 OF 15 HCAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2000:824294 HCAPLUS

DOCUMENT NUMBER: 133:359824

TITLE: Lawsonia-derived gene sodC and related

superoxide dismutase polypeptides, peptides and

proteins and their uses for diagnosis and treatment of avian and porcine infections

INVENTOR(S): Ankenbauer, Robert Gerard; Hasse, Detlef;

Panaccio, Michael; Rosey, Everett Lee; Wright,

Catherine

PATENT ASSIGNEE(S): Pfizer Products, Inc., USA; Pig Research and

Development Corp.; Agriculture Victoria Services

Pty., Ltd.

SOURCE: PCT Int. Appl., 85 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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APPLICATION NO. DATE
                   KIND DATE
    PATENT NO.
                    A1 20001123 WO 2000-AU436 20000511
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    WO 2000069903
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            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT,
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                     A1 20020206
                                        EP 2000-924975
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    JP 2003501013
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                                                          19990513
PRIORITY APPLN. INFO.:
                                                         20000511
                                      WO 2000-AU436
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The present invention relates generally to therapeutic compns. for the treatment and/or prophylaxis of intestinal disease conditions in animals and birds caused or exacerbated by Lawsonia intracellularis or similar or otherwise related microorganism. In particular, the present invention provides a novel gene derived from Lawsonia intracellularis which encodes an immunogenic SodC superoxide dismutase peptide,

polypeptide or protein that is particularly useful as an antigen in vaccine prepn. for conferring humoral immunity against Lawsonia intracellularis and related pathogens in animal hosts. The present invention is also directed to methods for the treatment and/or prophylaxis of such intestinal disease conditions and to diagnostic agents and procedures for detecting Lawsonia intracellularis or similar or otherwise related microorganisms.

REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 10 OF 15 HCAPLUS COPYRIGHT 2003 ACS L32000:588529 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 134:290822

Immunohistochemistry and polymerase chain TITLE: reaction for the detection of Lawsonia

intracellularis in porcine intestinal tissues with proliferative enteropathy

Kim, Junghyun; Choi, Changsun; Cho, Wan-Seob; AUTHOR(S):

Chae, Chanhee

Department of Veterinary Pathology, College of CORPORATE SOURCE:

Veterinary Medicine and School of Agricultural Biotechnology, Seoul National University, Suwon,

441-744, S. Korea

Journal of Veterinary Medical Science (2000), SOURCE:

62(7), 771-773

CODEN: JVMSEQ; ISSN: 0916-7250

Japanese Society of Veterinary Science PUBLISHER:

Journal DOCUMENT TYPE: English LANGUAGE:

Detection method of Lawsonia intracellularis was AB

studied in formalin-fixed paraffin-embedded intestinal tissues from 5 naturally infected pigs by immunohistochem. with a monoclonal

antibody against outer membrane protein

of L. intracellularis. Warthin-Starry silver

stain revealed clusters of argyrophilic, slightly curved rod-shaped organisms in the apical cytoplasm of enterocytes. Immunohistochem.

staining with a L. intracellularis-specific

monoclonal antibody confirmed the presence of the organism in the apical cytoplasm of hyperplastic enterocytes. The presence of

L. intracellularis in the ileum of pig with

proliferative enteropathy was confirmed by PCR further on the basis

of amplification of 319-bp products specific for porcine L . intracellularis chromosomal DNA. Immunohistochem. and

PCR may be a complementary method to confirm the diagnosis of

L. intracellularis infection in pigs.

THERE ARE 14 CITED REFERENCES AVAILABLE REFERENCE COUNT: 14

FOR THIS RECORD. ALL CITATIONS AVAILABLE

IN THE RE FORMAT

ANSWER 11 OF 15 HCAPLUS COPYRIGHT 2003 ACS L3

1997:494260 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 127:158904

Comparison of the 16S ribosomal DNA sequences TITLE: from the intracellular agents of proliferative enteritis in a hamster, deer, and ostrich with

the sequence of a porcine isolate of

Shears 308-4994 Searcher :

Lawsonia intracellularis

Cooper, Dale M.; Swanson, Debra L.; Barns, Susan AUTHOR(S):

M.; Gebhart, Connie J.

Division of Comparative Medicine, Research CORPORATE SOURCE:

Animal Resources, Medical School, University of

Minnesota, Minneapolis, MN, 55455, USA

SOURCE: International Journal of Systematic Bacteriology

(1997), 47(3), 635-639

CODEN: IJSBA8; ISSN: 0020-7713

American Society for Microbiology PUBLISHER:

DOCUMENT TYPE: Journal English LANGUAGE:

Proliferative enteritis is an enteric disease that affects a variety AB of animals. The causative agent in swine has been detd. to be an

obligate intracellular bacterium, Lawsonia

intracellularis, related to the sulfate-reducing bacterium Desulfovibrio desulfuricans. The intracellular agents found in the lesions of different animal species are antigenically

similar. In addn., strains from the pig, ferret, and hamster have been shown to be genetically similar. In this study we performed a partial 16S ribosomal DNA sequence anal. on the intracellular agent of proliferative enteritis from a hamster, a deer, and an ostrich

and compared these sequences to that of the porcine L. intracellularis isolate. Results of this study indicate

that the intracellular agents from these species with proliferative enteritis have high sequence similarity, indicating that they are all in the genus Lawsonia and that they may also be the same

species, L. intracellularis.

ANSWER 12 OF 15 HCAPLUS COPYRIGHT 2003 ACS L3

1997:459883 HCAPLUS ACCESSION NUMBER:

127:148036 DOCUMENT NUMBER:

Lymphoglandular complexes process TITLE:

antigen in the distal colon in swine

Mansfield, Linda S.; Hill, Doloros E.; Urban, AUTHOR(S):

Joseph F., Jr.

Department of Microbiology, College of CORPORATE SOURCE:

Veterinary Medicine, Michigan State University,

East Lansing, MI, 48824, USA

Cytokines, Cholera, and the Gut, [Papers from SOURCE:

the Joint Meeting of the United States-Japan Cooperative Medical Sciences Program Panels on Malnutrition and Cholera], Kiawah Island, S. C., Nov. 30-Dec. 3, 1995 (1997), Meeting Date 1995, 185-195. Editor(s): Keusch, Gerald T.; Kawakami, Masanobu. IOS Press: Amsterdam, Neth.

CODEN: 64SIAE

DOCUMENT TYPE: Conference; General Review

English LANGUAGE:

A review with 15 refs. We have identified a new antigen processing structure in the colon of swine that functions by recognizing and reacting to enteric bacteria that invade the host via the colon (I). Lymphoglandular complexes serve as antigen sampling structures and induce an immune response in exptl. infections of swine with whipworm. These studies grew from a clin. observation that swine with naturally acquired infections of whipworm had circumscribed nodular secondary bacterial lesions in the distal colon, distant from the site of worm attachment in the

> Shears 308-4994 Searcher :

proximal colon. Subsequent exptl. infections of weaned pigs revealed that low nos. of swine whipworm created an environment in the colon where opportunistic bacteria could invade and overgrow. Whipworm infected pigs developed severe diarrhea and failed to grow normally. Because enteric bacteria are known to utilize immune inductive sites for host invasion (2), we examd. this possibility in whipworm infected pigs. We found that bacteria selectively invaded and multiplied in the LGCs in specialized cells with M cell characteristics leading to expansion of the lymphoid cells in the underlying follicle. Both the worm and bacteria were necessary for these events to occur, and initiation of lesions was whipworm dose dependent. Propria nodules varied from 0.25 to 0.63 cm in diam. depending on the dose of whipworm. We isolated several species of bacteria from the LGC follicle deep to the muscularis mucosae. These included Campylobacter jejuni, C. coli, C. lari, and Escherichia coli. Lawsonia intracellularis was demonstrated in LGCs.

L3 ANSWER 13 OF 15 HCAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 1997:457165 HCAPLUS

DOCUMENT NUMBER: 12

127:94116

TITLE:

Lawsonia intracellularis

immunogenic components identification, DNA sequences, and uses for animal intestine

infection vaccine or diagnosis

INVENTOR(S):

Panaccio, Michael; Hasse, Detlef

PATENT ASSIGNEE(S):

Daratech Pty. Ltd., Australia; Pig Research and

Development Corporation; Panaccio, Michael;

Hasse, Detlef

SOURCE:

PCT Int. Appl., 94 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

1

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PA | TENT | NO. | | KI | ND | DATE | | | A | PPLI | CATI | ON NO | 0. | DATE | | |
|---------|------|---------|-----|-----|-------|------|------|-----|--------|------|------|-------|-----|------|------|-----|
| WO | 9720 | 050 | | | 1 | 1997 | 0605 | | _ ₩ | 0 19 | 96-A | U767 | | 1996 | 1129 | |
| | W: | AL, | AM, | AT, | AU, | AZ, | BA, | BB, | BG, | BR, | BY, | CA, | CH, | CN, | CU, | CZ, |
| • | | | | | | | | | | | | | | KG, | | |
| | | | | | | | | | | | | | | MW, | | |
| | | | | | | | | | | | | | | TR, | | |
| | | | | | | | | | | | | | | TM | • | |
| | RW: | | | | | | | | | | | | | | FR, | GB, |
| | | | | | | | | | | | | | | CI, | | |
| | | | | | | SN, | | | | • | • | • | · | · | • | • |
| CA | 2236 | | | | | | | | C | A 19 | 96-2 | 2365 | 74 | 1996 | 1129 | |
| | 9676 | | | | | | | | | U 19 | | | | 1996 | | |
| | 7183 | | | | | | | | | | | | | | | |
| | 8717 | | | | | | | | E | P 19 | 96-9 | 3886 | 3 | 1996 | 1129 | |
| | | | | | | | | | | | | | | NL, | | MC. |
| | • | | | | | LV, | | | , | , | , | , | | | , | • |
| CN | 1203 | 630 | , | Δ1, | , | 1998 | 1230 | | С | N 19 | 96-1 | 9866 | 6 | 1996 | 1129 | |
| | 9611 | | | | | | | | | R 19 | | | | 1996 | 1129 | |
| | 2000 | | | | | | | | _ | P 19 | | | | 1996 | 1129 | |
| | 3223 | | | | | | | | | | - | | - | 1996 | | |
| PRIORIT | | | | | | 2000 | | | AU 1 | | | | | 1995 | | |
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AU 1995-6911 A 19951130 WO 1996-AU767 W 19961129

The present invention relates generally to therapeutic compns. for AB the treatment and/or prophylaxis of intestinal disease conditions in animals and birds caused or exacerbated by Lawsonia intracellularis or similar or otherwise related microorganism. The present invention also contemplates methods for the treatment and/or prophylaxis of such intestinal disease conditions and to diagnostic agents and procedures for detecting Lawsonia intracellularis or similar or otherwise related microorganism. The Lawsonia intracellularis genomic library was screened with immunoscreened with anti-L. intracellularis sera. Clones found to be pos. according to immunoscreening were sequenced. GroEL and GroES proteins are two immunogenic components that were identified. Examples also included immunofluorescent detection of L. intracellularis bacteria in pig feces, formalin-killed vaccines, and putative vaccine candidate sequences.

L3 ANSWER 14 OF 15 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1995:405975 HCAPLUS

DOCUMENT NUMBER: 122:181343

TITLE: Intracellular domain of desmoglein 3 (pemphigus

vulgaris antigen) confers adhesive function on the extracellular domain of E-cadherin without binding catenins

AUTHOR(S): Roh, Joo-Young; Stanley, John R.

CORPORATE SOURCE: Natl. Cancer Inst., Natl. Inst. of Health,

Bethesda, MD, 20892, USA

SOURCE: Journal of Cell Biology (1995), 128(5), 939-47

CODEN: JCLBA3; ISSN: 0021-9525

PUBLISHER: Rockefeller University Press

DOCUMENT TYPE: Journal LANGUAGE: English

For the extracellular (EC) domain of E-cadherin to function in AR homophilic adhesion it is thought that its intracytoplasmic (IC) domain must bind .alpha.- and .beta.-catenins, which link it to the actin cytoskeleton. However, the IC domain of pemphigus vulgaris antigen (PVA or Dsg3), which is in the desmoglein subfamily of the cadherin gene superfamily, does not bind .alpha.- or .beta.-catenins. Because desmogleins have also been predicted to function in the cell adhesion of desmosomes, we speculated that the PVA IC domain might be able to act in a novel way in conferring adhesive function on the EC domain of cadherins. To test this hypothesis we studied aggregation of mouse fibroblast L cell clones that expressed chimeric cDNAs encoding the EC domain of E-cadherin with various IC domains. We show here that the full IC domain of PVA as well as an IC subdomain contg. only 40 amino acids of the PVA intracellular anchor (IA) region confer adhesive function on the E-cadherin EC domain without catenin-like assocns. with cytoplasmic mols. or fractionation with the cell cytoskeleton. This IA region subdomain is evolutionarily conserved in desmogleins, but not classical cadherins. These findings suggest an important cell biol. function for the IA region of desmogleins and demonstrate that strong cytoplasmic interactions are not absolutely necessarily for E-cadherin-mediated adhesion.

L3 ANSWER 15 OF 15 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1985:77080 HCAPLUS

DOCUMENT NUMBER: 102:77080

TITLE: Inherited deficiency of the Mac-1, LFA-1,

p150,95 glycoprotein family and its molecular

basis

AUTHOR(S): Springer, Timothy A.; Thompson, W. Scott;

Miller, Linda J.; Schmalstieg, Frank C.;

Anderson, Donald C.

CORPORATE SOURCE: Lab. Membrane Immunochem., Dana-Farber Cancer

Inst., Boston, MA, 02115, USA

SOURCE: Journal of Experimental Medicine (1984), 160(6),

1901-18

CODEN: JEMEAV; ISSN: 0022-1007

DOCUMENT TYPE:

Journal English

LANGUAGE: Leukocyte surface glycoproteins that share a common .beta. subunit AB have been found to be congenitally deficient in 3 unrelated patients with recurring bacterial infection. The glycoproteins, Mac-1, LFA-1, and p150,95 have the subunit compns. .alpha.M.beta., .alpha.L.beta., and .alpha.X.beta., resp. Using subunit-specific monoclonal antibodies, both the .alpha.M and .beta. subunits of Mac-1, the .alpha.L and .beta. subunits of LFA-1, and at the least the .beta. subunit of p150,95, were found to be deficient at the cell surface by the techniques of immunofluorescence flow cytometry, radioimmunoassay, and immunopptn. A latent pool of Mac-1 that can be expressed on granulocyte surfaces in response to secretory stimuli, such as formyl-Met-Leu-Phe, was also lacking in patients. Deficiency was found on all leukocytes tested, including granulocytes, monocytes, and T and B lymphocytes. Quantitation by immunofluorescence cytometry of subunits on granulocytes from parents of these patients and of a fourth deceased patient showed approx. half-normal surface expression, and, together with data on other siblings and a family with an affected father and children, demonstrate autosomal recessive inheritance. Deficiency appears to be quant. rather than qual., with 2 patients expressing .apprx.0.5% and 1 patient .apprx.5% of normal amts. The latter patient had .alpha..beta. complexes on the cell surface detectable by immunopptn. Biosynthesis expts. showed the presence of normal amts. of .alpha.'L intracellular precursor in lymphoid lines of all 3 patients. Together with surface deficiency of 3 mols. that share a common .beta. subunit but have differing .alpha. subnunits, this suggests the primary deficiencey is of the .beta. The lack of maturation of .alpha.'L to .alpha.L and the subunit. deficiency of the .alpha. subunits at the cell surface and in latent pools suggest that assocn. with the .beta. subunit is required for .alpha. subunit processing and transport to the cell surface or to latent pools. The mol. basis of this disease is discussed in light of adhesion-related functional abnormalities in patients' leukocytes and the blockade of similar functions in healthy cells by monoclonal antibodies.

(FILE 'MEDLINE, BIOSIS, EMBASE, WPIDS, CONFSCI, SCISEARCH, JICST-EPLUS, JAPIO, CABA, AGRICOLA, VETU, VETB' ENTERED AT 11:21:36 ON 10 APR 2003)

L4 9 S L1 L5 36 S L2

L6 44 S L4 OR L5

L7 21 DUP REM L6 (23 DUPLICATES REMOVED)

L7 ANSWER 1 OF 21 WPIDS (C) 2003 THOMSON DERWENT

ACCESSION NUMBER: 2002-557448 [59] WPIDS

DOC. NO. NON-CPI: N2002-441304 DOC. NO. CPI: C2002-158153

TITLE: New immunogenic polypeptide comprising epitope of

Lawsonia spp. polypeptide such as fihB, fliR, ntrC, glnH, motA, polypeptides, useful in vaccines for treatment of porcine proliferative enteropathy in

pigs and birds.

DERWENT CLASS: B04 C06 D16 S03

INVENTOR(S): GOOD, R T; KING, K W; ROSEY, E L; STRUGNELL, R A
PATENT ASSIGNEE(S): (AGRI-N) AGRIC VICTORIA SERVICES PTY LTD; (AUPO-N)

AUSTRALIAN PORK LTD; (PFIZ) PFIZER PROD INC

COUNTRY COUNT: 98

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2002038594 A1 20020516 (200259)* EN 155

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ

NO NZ OM PH PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA

UG US UZ VN YU ZA ZW

AU 2002014810 A 20020521 (200260)

APPLICATION DETAILS:

| PATENT NO K | IND | APE | PLICATION | DATE |
|--------------------------------|-----|-----|--------------|----------------------|
| WO 2002038594 AU 2002014810 | | | 2002 1102102 | 20011109 20011109 |

FILING DETAILS:

| PATENT NO | KIND | PATENT NO |
|-------------|---------------|--------------|
| | | |
| AU 20020148 | 10 A Based on | WO 200238594 |

PRIORITY APPLN. INFO: US 2000-249596P 20001117; AU 2000-1381 20001110

AN 2002-557448 [59] WPIDS

AB WO 200238594 A UPAB: 20020916

NOVELTY - An isolated or recombinant immunogenic polypeptide (I) which comprises, mimics or cross-reacts with a B-cell or T-cell epitope of a Lawsonia spp. polypeptide such as fihB, fliR, ntrC, glnH, motA, motB, tlyC, ytfM or ytfN polypeptides, is new.

DETAILED DESCRIPTION - An isolated or recombinant immunogenic polypeptide (I) which comprises, mimics or cross-reacts with a B-cell or T-cell epitope of a Lawsonia spp. polypeptide such as fihB, fliR, ntrC, glnH, motA, motB, tlyC, ytfM or ytfN polypeptides, is:

(i) a polypeptide of Lawsonia spp. which comprises an amino acid sequence that has at least about 60% sequence identity overall

- to a fully defined amino acid (PS) sequence of 207 (S2), 262 (S4), 456 (S6), 137 (S8), 282 (S10), 237 (S12), 348 (S14), 602 (S16), or 1382 (S18) amino acids as given in specification;
- (ii) a polypeptide of Lawsonia spp. which comprises an amino acid sequence which has at least 60% sequence identity overall to an amino acid sequence encoded by L. intracellularis
- (Li) DNA contained within a plasmid (P) having AGAL Accession Nos: NM00/16476 (plasmid pGTE1 glnH); NM00/16477 (plasmid pGTE2 flhB);
- NM00/16478 (plasmid pGTE3 fliR); NM00/16479 (plasmid pGTE4 motA/B);
- NM00/16480 (plasmid pGTE5 tlyC); NM00/16481 (plasmid pGTE6 ntrC);
- NM00/16482 (plasmid pGTE7 ytfM); or NM01/23286 (plasmid pGTE8 ytfN);
- (iii) a polypeptide which comprises at least about 5 contiguous amino acids of PS;
- (iv) a polypeptide which comprises at least about 5 contiguous amino acids of amino acid sequence of Li DNA contained within (P);
- (v) a polypeptide which comprises an amino acid sequence encoded by nucleotide sequence of Lawsonia spp. having at least 60% identity overall to a fully defined nucleotide sequence (NS) of 622 (S1), 789 (S3), 1371 (S5), 412 (S7), 849 (S9), 717 (S11), 1047 (S13), 1812 (S15), or 4149 (S17) nucleotides as given in specification;
- (vi) a polypeptide which comprises an amino acid sequence encoded by a nucleotide sequence of Lawsonia spp. having at least 60% sequence identity overall to nucleotide sequence of Li DNA contained with an (P);
- (vii) a polypeptide encoded by at least 15 contiguous nucleotides of NS;
- (viii) a polypeptide encoded by at least 15 contiguous nucleotides of nucleotide sequence of Li DNA contained within (P);
- (ix) a homolog, analog or derivative of above mentioned polypeptides which mimic a B-cell or T-cell epitope of Lawsonia spp. INDEPENDENT CLAIMS are also included for the following:
- (1) a vaccine composition (II) for the prophylaxis or treatment of infection of an animal by Lawsonia spp. which comprises an immunogenic component that comprises (I) and one or more carriers, diluents or adjuvants suitable for veterinary or pharmaceutical use;
- (2) a combination vaccine composition (III) for the prophylaxis or treatment of infection of an animal by Lawsonia spp., comprising:
 - (i) a first immunogenic component which comprises (I); and
- (ii) a second immunogenic component different from first immunogenic component and comprising a Li polypeptide such as FlgE, hemolysin, OmpH, SodC, flhB, fliR, ntrC, glnH, motA, motB, tlyC, ytfM, or ytfN polypeptides and one or more carriers, diluents or adjuvants suitable for veterinary or pharmaceutical use;
- (3) a vaccine vector (IV) that comprises, in an expressible form, an isolated nucleic acid molecule (V) comprising a nucleotide sequence such as:
- (i) a protein-encoding nucleotide sequence having at least 60% sequence identity overall to a sequence of NS;
- (ii) a protein-encoding nucleotide sequence having at least 60% identity overall to the protein-encoding sequence of Li DNA contained within (P);
- (iii) a protein-encoding nucleotide sequence which comprises at least about 15 contiguous nucleotides of NS;
- (iv) a protein-encoding nucleotide sequence which comprises at least 15 contiguous nucleotides of Li DNA contained within (P);
 - (v) a protein-encoding nucleotide sequence which hybridizes

under low stringency condition to the complement of NS;

- (vi) a protein-encoding nucleotide sequence which hybridizes under low stringency conditions to non-coding strand of Li DNA contained within (P); and
- (vii) a homolog, analog or derivative of above mentioned nucleotide sequences which encodes the polypeptide that mimics a B-cell or T-cell epitope of Lawsonia spp.;
- (4) an isolated polyclonal or monoclonal antibody molecule (VI) that binds specifically to Lawsonia spp. polypeptide of flhB, fliR, ntrC, glnH, motA, motB, tlyC, ytfM, or ytfN polypeptide, or homolog, analog or derivative of the above mentioned polypeptide;
- (5) an isolated nucleic acid molecule (N) which consists of a nucleotide sequence encoding Lawsonia spp. such as flhB, fliR, ntrC, glnH, motA, motB, tlyC, ytfM, or ytfN;
- (6) a probe or primer comprising any one of fully defined 50 oligonucleotide sequences as given in specification such as catattcaaggtacagcatctgatgg, ctcctttacaaaccttgctcc, gctcatctaaagaacactttcc, caaggtagtatacaacttattgg, etc., or complementary nucleotide sequence to the oligonucleotide sequence;
- (7) a plasmid having AGAL Accession Nos: NM00/16476 (plasmid pGTE1 glnH); NM00/16477 (plasmid pGTE2 flhB); NM00/16478 (plasmid pGTE3 fliR); NM00/16479 (plasmid pGTE4 motA/B); NM00/16480 (plasmid pGTE5 tlyC); NM00/16481 (plasmid pGTE6 ntrC); NM00/16482 (plasmid pGTE7 ytfM); or NM01/23286 (plasmid pGTE8 ytfN);
- (8) a recombinant vector (VII) capable of replication in a host cell, where the vector comprises (N);
 - (9) a host cell (VIII) comprising (VII);
- (10) identifying (M1) whether or not a porcine or avian animal has suffered from a past infection, or is currently infected, with Li or a microorganism that is immunologically cross-reactive with Li;
- (11) diagnosing (M2) infection of a porcine or avian animal by Li or a microorganism that is immunologically cross-reactive with Li; and
- (12) detecting (M3) Li or related microorganism in a biological sample derived from a porcine or avian animal subject.

ACTIVITY - Antibacterial.

MECHANISM OF ACTION - Vaccine. No supporting data is given.

USE - (I) is useful for identifying whether or not a porcine or avian animal has suffered from a past infection, or is currently infected, with Li or a microorganism that is immunologically cross-reactive with Li. (VI) is useful for diagnosing infection of a porcine or avian animal by Li or a microorganism that is immunologically cross-reactive with Li. (N) is useful as probes or primers for detecting Li or related microorganism in a biological sample derived from a porcine or avian animal subject (all claimed). (I) is preferably useful for vaccinating porcine animals against porcine proliferative enteropathy (PPE). (I) is also useful in vaccines for the prophylaxis and treatment of PPE in birds. (II) is useful for conferring protection against infection by other species of the genus Lawsonia or other microorganisms related to Li.

Dwg.0/1

L7 ANSWER 2 OF 21 WPIDS (C) 2003 THOMSON DERWENT ACCESSION NUMBER: 2002-519087 [55] WPIDS

DOC. NO. CPI:

C2002-146759

TITLE:

A proliferative ileitis vaccine useful for protecting mammals from disease caused by

Lawsonia intracellularis,

comprises tissue culture grown Lawsonia

intracellularis.

DERWENT CLASS:

A96 B04 C06 D16

PATENT ASSIGNEE(S):

(ARIZ-N) ARIZONA BOARD OF REGENTS

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2002026250 A2.20020404 (200255)* EN 43

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK

DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP

KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT

RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU ZA ZW

AU 2001093151 A 20020408 (200255)

92

APPLICATION DETAILS:

| PATENT NO KIND | APPLICATION | DATE |
|------------------|-----------------|----------|
| | | |
| WO 2002026250 A2 | WO 2001-US30284 | 20010927 |
| AU 2001093151 A | AU 2001-93151 | 20010927 |

FILING DETAILS:

| PATENT NO | KIND | | PAT | ENT NO |
|-------------|------|----------|-----|-----------|
| | | | | |
| AU 20010931 | 51 A | Based on | WO | 200226250 |

PRIORITY APPLN. INFO: US 2000-677108 20000929

AN 2002-519087 [55] WPIDS

AB WO 200226250 A UPAB: 20020829

NOVELTY - A proliferative ileitis vaccine (I), comprising tissue culture grown Lawsonia intracellularis (II),

which produces antibodies in pigs reacting with at least one of the antigens (A) selected from 21 kDa, 31 kDa, 41 kDa, 44 kDa,

60 kDa, 71 kDa, 115 kDa and greater than 115 kDa, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) a monoclonal antibody recognizing (II) antigen with a molecular weight of (A);
- (2) growing (II) in a susceptible tissue culture to protect a mammal against proliferative ileitis caused by (II);
 - (3) producing a proliferative ileitis vaccine, by:
- (a) growing (II) in a susceptible tissue culture, harvesting the tissue culture grown (II), inactivating or stabilizing the harvest and adjuvanting the inactivated harvest into a vaccine or formulating the stabilized harvest to produce a vaccine;
- (b) growing and harvesting (II) as above, inactivating the harvest, extracting a protective **antigen** from the harvested inactivated tissue culture grown to produce a subunit and adjuvanting the subunit to produce a vaccine;
- (c) identifying a target immunogen of (II), constructing and screening (II) genomic library, identifying the recombinant clone, producing the target immunogen of (II), identifying a gene encoding

an immunoreactive group using the production vector and formulating the immunoreactive group into a vaccine;

- (d) identifying a gene encoding an immunoreactive group of a target immunogen of (II) as above, expressing the immunoreactive group using a live production vector and formulating the vector into a vaccine;
- (e) identifying a target immunogen of (II), sequencing the target immunogen of (II), inserting the sequence of the target immunogen into a production vector, expressing the target immunogen by the production vector, growing the production vector expressing the target immunogen and formulating the target immunogen into a vaccine; or
- (f) preparing a monoclonal antibody to a functional immunogen of (II), identifying a functional immunogen detected by the monoclonal antibody as a target immunogen, sequencing the immunogen, expressing the target immunogen in a production vector, growing the production vector to express a target immunogen, and formulating the target immunogen into a vaccine;
 - (4) producing a proliferative ileitis subunit vaccine;
- (5) diagnosing proliferative ileitis, by detecting a target immunogen of (II) by an assay such as a fluorescence assay (FA), immunofluorescence assay (IFA), polymerase chain reaction (PCR) and enzyme linked immunosorbent assay (ELISA); and
- (6) quantitating **antigenic** mass during **antigen** production, by detecting an **antigen** the molecular weight of (A).

ACTIVITY - Antibacterial; Antiinflammatory. MECHANISM OF ACTION - Vaccine (claimed).

To determine whether (I) could protect pigs from a homologous challenge or from exposure to heterologous isolates or strains, ten 4-week-old pigs were vaccinated and later challenged. Ten control pigs received equal doses of a mock vaccine which contained only the tissue culture, medium Minimal Essential Medium (MEM) and adjuvant (without antigen). The pigs were numbered and then placed into two different treatment groups to provide two different repetitions of each treatment. The pigs were challenged to (II) through incubation with 75ml of viable (II)-infected cells per pig (5 days post cell-culture infection) 21 days after the booster vaccination. Pigs were observed for clinical signs of disease for 24 days and then necropsied and examined for lesions of ileitis (gross lesions and hyperplasia). Rectal swabs were cultured for S.hyodysenteriae and Salmonella spp. prior to challenge and at necropsy. None of these swabs were positive indicating that pigs were not infected with S.hyodysenteriae or Salmonella spp.. Two pigs (one vaccinated and one control) died of respiratory lesions prior to challenge. The remaining control pigs showed sporadic diarrhea. None of the vaccinated pigs exhibited any grossly observable pathology. Upon necropsy at 24 days following challenge, seven of eight vaccinated pigs were normal, whereas, five of nine control pigs had gut lesions typical of (II). One vaccinate had both gross lesions and hyperplasia, whereas, five control pigs showed both gross lesions and hyperplasia. One control had hyperplasia but showed no gross lesions.

USE - (I) is useful for protecting a mammal through vaccination, from a disease caused by (II), especially proliferative ileitis (claimed). Dwg.0/6

L7 ANSWER 3 OF 21 WPIDS (C) 2003 THOMSON DERWENT

ACCESSION NUMBER: 2002-521947 [56] WPIDS

DOC. NO. NON-CPI: N2002-413067 DOC. NO. CPI: C2002-147814

TITLE: New Lawsonia intracellularis

proteins, useful as a vaccine or for manufacturing

a vaccine for combating L.

intracellularis infections, e.g. porcine

proliferative enteropathy, which is an important

disease in the pig industry.

DERWENT CLASS: B04 C04 D16 S03

INVENTOR(S): JACOBS, A A C; VERMEIJ, P
PATENT ASSIGNEE(S): (ALKU) AKZO NOBEL NV

COUNTRY COUNT: 30

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

EP 1219711 A2 20020703 (200256)* EN 26

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI TR

AU 2001097371 A 20020627 (200256)

CA 2365494 A1 20020620 (200256) EN

JP 2003000276 A 20030107 (200314) 71

HU 2001005379 A2 20030128 (200323)

APPLICATION DETAILS:

| PATENT NO K | IND | API | PLICATION | DATE |
|---------------|-----|-----|--------------|----------|
| EP 1219711 | A2 | EP | 2001-204919 | 20011214 |
| AU 2001097371 | A | ΑU | 2001-97371 | 20011220 |
| CA 2365494 | A1 | CA | 2001-2365494 | 20011218 |
| JP 2003000276 | A | JP | 2001-385373 | 20011219 |
| HU 2001005379 | A2 | HU | 2001-5379 | 20011219 |
| | | | | |

PRIORITY APPLN. INFO: EP 2000-204660 20001220

AN 2002-521947 [56] WPIDS AB EP 1219711 A UPAB: 20020903

NOVELTY - Lawsonia intracellularis proteins (I)

comprising a fully defined sequence at least 70% homologous to the sequence comprising 218 amino acids (P1) or 475 amino acids (P2) given in the specification, or their immunogenic fragments, are new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included

for:

- (1) nucleic acid sequences encoding the L. intracellularis proteins (or a part of the nucleic acid sequence that encodes an immunogenic fragment of the proteins) comprising a sequence with at least 70% homology with the nucleic acid sequence having 656 bp (NA1) or 1428 bp (NA2) fully defined in the specification;
- (2) deoxyribonucleic acid (DNA) fragment comprising the nucleic acid;
- (3) a recombinant DNA molecule comprising the nucleic acid sequences above, or the DNA fragment, under the control of a functionally linked promoter;
 - (4) a live recombinant carrier comprising the DNA fragment or

the recombinant DNA molecule;

(5) a host cell comprising the NA1 or NA2 nucleic acid sequences, the DNA fragment, the recombinant DNA molecule or the live recombinant carrier;

L. intracellularis Outer

Membrane Protein, which has a molecular weight of 19.21 kD, or its immunogenic fragment, obtainable by a process comprising:

- (a) subjecting an outer membrane preparation to sodium dodecyl sulfate polyacrylamide gel electrophoresis (SDS-PAGE); and
 - (b) excision of the 19 or 21 kD band from the gel;
 - (6) a vaccine for combating L.

intracellularis infections comprising the NA1 or NA2 nucleic
acid sequences, the DNA fragment, the recombinant DNA molecule, the
live recombinant carrier, the host cell, or the P1 or P2 L
. intracellularis proteins; and a pharmaceutical carrier;

- (7) preparing the vaccine by admixing the NA1 or NA2 nucleic acid sequences, the DNA fragment, the recombinant DNA molecule, the live recombinant carrier, the host cell, or the P1 or P2 L . intracellularis proteins; and a pharmaceutical carrier; and
- (8) a diagnostic test for detecting a L. intracellularis DNA comprising the NA1 or NA2 nucleic acid sequences, or a fragment of these sequences with a length of at least 12, preferably 18, nucleotides.

ACTIVITY - Antibiotic. No suitable data given. MECHANISM OF ACTION - Vaccine.

USE - (I) are useful as a vaccine or for manufacturing a vaccine for combating L. intracellularis infections (claimed), e.g. porcine proliferative enteropathy, which an important disease in the pig industry. (I) is also useful for diagnosing L. intracellularis infection and for detecting L. intracellularis DNA, L. intracellularis antigens or antibodies against L. intracellularis.

Dwg.0/2

L7 ANSWER 4 OF 21 MEDLINE DUPLICATE 1

ACCESSION NUMBER: 2002284767 MEDLINE

DOCUMENT NUMBER: 22006891 PubMed ID: 12010978
TITLE: LsaA, an antigen involved in cell

attachment and invasion, is expressed by

Lawsonia intracellularis during

infection in vitro and in vivo.

AUTHOR: McCluskey Jackie; Hannigan Joanne; Harris Jennifer D;

Wren Brendan; Smith David G E

CORPORATE SOURCE: Zoonotic & Animal Pathogens Research Laboratory,

Department of Medical Microbiology, Easter Bush

Veterinary Centre, University of Edinburgh,

Edinburgh, United Kingdom.

SOURCE: INFECTION AND IMMUNITY, (2002 Jun) 70 (6) 2899-907.

Journal code: 0246127. ISSN: 0019-9567.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals
OTHER SOURCE: GENBANK-AF498259

ENTRY MONTH: 200206

ENTRY DATE: Entered STN: 20020528

Last Updated on STN: 20020627 Entered Medline: 20020626

AB Lawsonia intracellularis has been identified

recently as the etiological agent of proliferative enteropathies, which are characterized by intestinal epithelial hyperplasia and associated moderate immune responses. This disease complex has been reported in a broad range of animals, prevalently in pigs, and

L. intracellularis has been linked with ulcerative

colitis in humans. L. intracellularis is an

obligate intracellular bacterium, and the pathogenic mechanisms used to cause disease are unknown. Using in vitro-grown organisms as a source of genomic DNA, we identified a Lawsonia gene which encodes a

surface antigen, LsaA (for Lawsonia surface

antigen), associated with attachment to and entry into
cells. The deduced amino acid sequence of this protein showed some
similarity to members of a novel protein family identified in a
number of other bacterial pathogens but for which roles are not
fully defined. Transcription of this gene was detected by reverse
transcription-PCR in L. intracellularis grown in

vitro in IEC18 cells and in bacteria present in ileal tissue from infected animals. Immunohistochemistry with specific monoclonal antibody and immunoblotting with sera from infected animals

demonstrated that LsaA protein is synthesized by L.

intracellularis during infection. Expression of this gene
during infection in vitro and in vivo suggests that this surface
antigen is involved during infection, and phenotypic

analysis indicated a role during L. intracellularis attachment to and entry into intestinal

epithelial cells

L7 ANSWER 5 OF 21 MEDLINE DUPLICATE 2

ACCESSION NUMBER: 2002486170 IN-PROCESS DOCUMENT NUMBER: 22232420 PubMed ID: 12296397

TITLE: A comparative study of an indirect fluorescent

antibody test and an immunoperoxidase monolayer assay

for the diagnosis of porcine proliferative

enteropathy.

AUTHOR: Guedes Roberto M C; Gebhart Connie J; Winkelman

Nathan L; Mackie-Nuss Rebecca A

CORPORATE SOURCE: Department of Veterinary PathoBiology, College of

Veterinary Medicine, University of Minnesota, Saint

Paul 55108, USA.

SOURCE: JOURNAL OF VETERINARY DIAGNOSTIC INVESTIGATION, (2002

Sep) 14 (5) 420-3.

Journal code: 9011490. ISSN: 1040-6387.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: English
FILE SEGMENT: IN-PROCESS; NONINDEXED; Priority Journals

ENTRY DATE: Entered STN: 20020926

Last Updated on STN: 20021213

The currently used indirect fluorescent antibody test (IFAT) for the detection of antibodies against porcine proliferative enteropathy (PPE) was compared to an immunoperoxidase monolayer assay (IPMA). Serum samples used in this comparison were collected from 5-week-old pigs on day 0 (pre-experimental challenge) and on days 7, 14, 21,

and 28 after oral inoculation with intestinal homogenate from pigs affected by PPE (28 challenged pigs) and sucrose phosphate glutamate solution (2 control pigs). All animals were euthanized 4 weeks after inoculation. Immunohistochemistry staining was applied to formalin-fixed, paraffin-embedded sections of ileum for the detection of Lawsonia intracellularis antigen. The serology results with each method agreed in all samples, except on days 0 and 7 in 1 control animal, which was positive by IPMA, but negative by IFAT. The percentage of agreement between IFAT and IPMA was 98.6%.

L7 ANSWER 6 OF 21 MEDLINE . DUPLICATE 3

ACCESSION NUMBER: 2001263511 MEDLINE

DOCUMENT NUMBER: 21254310 PubMed ID: 11355669

TITLE: Granulomatous enteritis and lymphadenitis in Iberian

pigs naturally infected with Lawsonia

intracellularis.

AUTHOR: Segales J; Fernandez-Salguero J M; Fructuoso G;

Quintana J; Rosell C; Pozo J; De Arriba M L; Rubio P;

Domingo M

CORPORATE SOURCE: Department de Sanitat i Anatomia Animals, Facultat de

Veterinaria, Universitat Autonoma de Barcelona,

Spain.. Joaquim.Segales@uab.es

SOURCE: VETERINARY PATHOLOGY, (2001 May) 38 (3) 343-6.

Journal code: 0312020. ISSN: 0300-9858.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200110

ENTRY DATE: Entered STN: 20011015

Last Updated on STN: 20011015 Entered Medline: 20011011

AΒ Intestinal samples and/or lymph nodes of two Iberian pigs from two different farms were submitted for histopathologic examination. Both pigs had proliferation of ileal and/or cecal crypts with almost complete absence of goblet cells. Infection by Lawsonia intracellularis was demonstrated by immunohistochemistry and polymerase chain reaction assay. The mesenteric lymph node of one pig had moderate lymphocyte depletion with granulomatous inflammation of the lymph node parenchyma. Histiocytes and multinucleated giant cells from the lymph node of one pig contained L. intracellularis antigen within the cytoplasm. This pig had also porcine circovirus type 2 (PCV-2) infection, but nucleic acid and antigen of this virus were not demonstrated in the lymph node. The second pig had lymphocyte depletion and marked granulomatous inflammation in Peyer's patches. Histiocytes and multinucleated giant cells in areas of granulomatous inflammation contained L. intracellularis antigen; no PCV-2 nucleic acid or antigen was detected in the tissues of this pig. This is the first description of granulomatous ileitis and lymphadenitis associated with ${f L}$

L7 ANSWER 7 OF 21 WPIDS (C) 2003 THOMSON DERWENT

ACCESSION NUMBER: 2001-016212 [02] WPIDS

DOC. NO. CPI: C2001-004517

. intracellularis infection.

TITLE: New immunogenic Lawsonia hemolysin peptide, nucleic

acid and antibody, useful in vaccines and for the diagnosis of Lawsonia infections, especially in

swine.

DERWENT CLASS:

B04 D16

INVENTOR(S):
PATENT ASSIGNEE(S):

ANKENBAUER, R G; HASSE, D; PANACCIO, M; ROSEY, E L
(AGRI-N) AGRIC VICTORIA SERVICES PTY LTD; (PFIZ)

DELGER PROD INC. (BICE-N) BIC RES (DEV CORR.

PFIZER PROD INC; (PIGR-N) PIG RES & DEV CORP;

(AUPO-N) AUSTRALIAN PORK LTD

COUNTRY COUNT:

93

PATENT INFORMATION:

| PATENT NO | KIND DATE | WEEK | LA | PG |
|-----------|-----------|------|----|----|
| | | | | |

WO 2000069906 A1 20001123 (200102)* EN 95

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA

RO RO SD SE SG SI SK SE TJ TM TK TT 12 OA OG OS OZ VN 10 ZA

zw

AU 2000043861 A 20001205 (200113)

EP 1177213 A1 20020206 (200218) EN

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI

APPLICATION DETAILS:

| PATENT NO K | IND | API | PLICATION | DATE |
|--|-----|----------|---|--|
| WO 2000069906 AU 2000043861 EP 1177213 | | AU EP | 2000-AU439 2000-43861 2000-924978 2000-AU439 | 20000511 20000511 20000511 20000511 |

FILING DETAILS:

| PAT | TENT NO K | IND | | | PAT | TENT NO |
|-----|------------|--------------|-------|-----|-----|------------|
| 711 | 2000043861 | | Pacod | | พด | 200069906 |
| | 1177213 | | Based | | | 200069906 |
| E E | 11//213 | Δ_{T} | Dasea | OII | " | 2000000000 |

PRIORITY APPLN. INFO: US 1999-134022P 19990513

AN 2001-016212 [02] WPIDS

AB WO 200069906 A UPAB: 20010110

NOVELTY - Isolated or recombinant polypeptide (I) that comprises, mimics or cross-reacts with a B- or T-cell epitope of a hemolysin polypeptide from a Lawsonia spp.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) a vaccine comprising, at least one carrier, diluent or adjuvant and a (I) having at least 70% sequence identity with a fully defined 251 aa sequence (1), (given in the specification), or at least 50% identity overall with aa 1-50 of (1), or their immunogenic homolog, analog or derivative that is immunologically cross-reactive with L. intracellularis;
- (2) vaccine vector comprising a nucleic acid sequence (II) that encodes (1);

- (3) poly- or monoclonal antibody (Ab) that binds to Lawsonia hemolysin polypeptide, or its derivatives, that have at least 70% sequence identity with (1);
- (4) an isolated nucleic acid (III) that encodes a peptide, oligopeptide or polypeptide having at least 70% sequence identity with (1), at least 50% identity overall with aa 1-50 of (1), or its homolog, analog or derivative that mimics a B- or T-cell epitope, also complements of (III);
- (5) a probe or primer containing at least 15 contiguous nucleotides from a 756 bp sequence (2), reproduced, or its complement; and
 - (6) the plasmid pALK12 (ATCC 207195).

ACTIVITY - Antibacterial.

MECHANISM OF ACTION - Induction of a specific humoral immune response.

USE - (I) are used (i) as antigens in vaccines to prevent or treat infection by Lawsonia, in birds and animals, especially pigs, to raise specific antibodies (Ab) and to detect past or present infection. Ab are also useful in diagnosis, to detect L. intracellularis or immunologically cross-reactive species, also for identification of epitopes in hemolysin. Vectors that contain nucleic acid (II) that encodes (I) are also useful in genetic vaccines, and fragments of (II) are useful as primers or probes for detecting L. intracellularis or related microorganisms, in hybridization or amplification assays.

Dwg.0/1

L7 ANSWER 8 OF 21 WPIDS (C) 2003 THOMSON DERWENT

ACCESSION NUMBER:

2001-016211 [02] WPIDS

DOC. NO. CPI:

C2001-004516

TITLE:

New isolated Lawsonia spp. OmpH polypeptides and

nucleic acids, useful for the prophylaxis,

treatment and detection of Lawsonia infections.

DERWENT CLASS:

B04 D16

INVENTOR(S):

HASSE, D; PANACCIO, M; SINISTAJ, M

PATENT ASSIGNEE(S):

(AGRI-N) AGRIC VICTORIA SERVICES PTY LTD; (PIGR-N) PIG RES & DEV CORP; (AUPO-N) AUSTRALIAN PORK LTD

93

COUNTRY COUNT: 9

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2000069905 A1 20001123 (200102) * EN 84

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA

AU 2000043860 A 20001205 (200113)

EP 1183268 A1 20020306 (200224) EN

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI

BR 2000011290 A 20020521 (200238)

APPLICATION DETAILS:

| PATENT NO KIND | APPLICATION | DATE |
|-------------------------------------|---------------------------------|----------------------|
| WO 2000069905 A1 AU 2000043860 A | WO 2000-AU438 AU 2000-43860 | 20000511 |
| EP 1183268 A1 | EP 2000-924977 WO 2000-AU438 | 20000511 20000511 |
| BR 2000011290 A | BR 2000-11290 WO 2000-AU438 | 20000511 20000511 |

FILING DETAILS:

| PAI | ENT NO K | IND | | | PAT | CENT NO | |
|-----|------------|-----|-------|----|-----|-----------|--|
| ĀU | 2000043860 | A | Based | on | WO | 200069905 | |
| ΕP | 1183268 | A1 | Based | on | WO | 200069905 | |
| BR | 2000011290 | Α | Based | on | WΟ | 200069905 | |

PRIORITY APPLN. INFO: US 1999-133986P 19990513

AN 2001-016211 [02] WPIDS

AB WO 200069905 A UPAB: 20010110

NOVELTY - A novel isolated or recombinant immunogenic polypeptide mimics or cross-reacts with a B-cell or T-cell epitope of a Lawsonia spp. OmpH polypeptide.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) an isolated or recombinant immunogenic polypeptide comprising:
- (i) a peptide, oligopeptide or polypeptide which comprises an amino acid sequence having at least about 70% sequence identity overall to a fully defined 186 aa sequence (I) (given in the specification); or
- (ii) a homolog, analog or derivative of (i) which mimics a B-cell or T-cell epitope of a Lawsonia spp. OmpH polypeptide;
- (2) a vaccine composition for the prophylaxis or treatment of infection of an animal by Lawsonia spp., comprising an immunogenic component derived from an isolated or recombinant polypeptide having at least about 70% sequence identity overall to (I) or an immunogenic homolog, analog or derivative which is immunologically cross-reactive with L. intracellularis, and one or more carriers, diluents or adjuvants;
- (3) a combination vaccine composition for the prophylaxis or treatment of infection of an animal by Lawsonia spp .comprising:
- (i) a first immunogenic component comprising an isolated or recombinant polypeptide having at least about 70% sequence identity to (I) or an immunogenic homolog, analog, or derivative which is immunologically cross-reactive with L. intracellularis;
- (ii) a second immunogenic component comprising an antigenic L. intracellularis peptide, polypeptide or protein; and
- (iii) one or more carriers, diluents or adjuvants suitable for veterinary or pharmaceutical use;
- (4) a vaccine vector that comprises, in an expressible form, an isolated nucleic acid molecule having a nucleotide sequence that encodes (I), such that the immunogenic polypeptide is expressible at a level to confer immunity against Lawsonia spp., when administered to a porcine or avian animal;

- (5) a poly- or monoclonal antibody molecule capable of binding specifically to a OmpH polypeptide or a derivative of a OmpH polypeptide that is derived from Lawsonia spp. having at least about 70% sequence identity to (I);
- (6) an isolated nucleic acid molecule (NAM) comprising a sequence of nucleotides, or their complements which encode, a peptide, oligopeptide or polypeptide selected from:
- (i) a peptide, oligopeptide or polypeptide which comprises an amino acid sequence which has at least about 70% sequence identity overall to an amino acid sequence (I); and
- (ii) a homolog, analog or derivative of (i) which mimics a B-cell or T-cell epitope of Lawsonia spp.;
- (7) a method of detecting L. intracellularis or related microorganism in a biological sample derived from a porcine or avian animal subject comprising hybridizing one or more probes or primers derived from a fully defined 561 bp nucleotide sequence (NS) (II), or its complements to the sample and then detecting the hybridization using a detection device;
- (8) a probe or primer having at least about 15 contiguous nucleotides in length derived from (II) or its complements;
 - (9) a plasmid designated pALK13 (ATCC No: 207196).

USE - The polypeptides are capable of eliciting the production of antibodies against Lawsonia spp. when administered to an avian or porcine animal (claimed). They can be used for conferring a protective immune response against Lawsonia spp. when administered to an avian or porcine animal (claimed). They can be used for the prophylaxis or treatment of an infection of an animal by Lawsonia spp. (claimed). The nucleic acids can also be used for prophylaxis or treatment of infections. The products can also be used for detection, e.g. for detecting whether or not a porcine or avian animal has suffered from a past infection or is currently infected with L. intracellularis. They are used particularly for porcine proliferative enteropathy (PPE) infections. Dwg.0/3

ANSWER 9 OF 21 WPIDS (C) 2003 THOMSON DERWENT WPIDS

ACCESSION NUMBER:

2001-016210 [02]

DOC. NO. CPI:

C2001-004515

TITLE:

New immunogenic Lawsonia FlgE peptide, its nucleic acid and antibody, useful in vaccines and diagnosis of Lawsonia infections, particularly in swine.

DERWENT CLASS:

B04 D16

INVENTOR(S):

ANKENBAUER, R G; HASSE, D; PANACCIO, M; PARSONS, J;

ROZEY, E L; SINISTAJ, M; ROSEY, E L

PATENT ASSIGNEE(S):

(AGRI-N) AGRIC VICTORIA SERVICES PTY LTD; (PFIZ) PFIZER PROD INC; (PIGR-N) PIG RES & DEV CORP;

(AUPO-N) AUSTRALIAN PORK LTD

COUNTRY COUNT:

93

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG _____

WO 2000069904 A1 20001123 (200102)* EN 95

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT

RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2000043859 A 20001205 (200113)

EP 1181315 A1 20020227 (200222) EN

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI

BR 2000011294 A 20020226 (200223)

APPLICATION DETAILS:

| PATENT NO K | IND | API | PLICATION | DATE |
|---------------|-----|-----|-------------|----------|
| | | | | |
| WO 2000069904 | A1 | WO | 2000-AU437 | 20000511 |
| AU 2000043859 | A | ΑU | 2000-43859 | 20000511 |
| EP 1181315 | A1 | ΕP | 2000-924976 | 20000511 |
| | | WO | 2000-AU437 | 20000511 |
| BR 2000011294 | A | BR | 2000-11294 | 20000511 |
| | • | WO | 2000-AU437 | 20000511 |

FILING DETAILS:

| PAT | TENT NO I | KIND | | | PA' | TENT NO |
|-----|------------|------|-------|----|-----|-----------|
| AU | 2000043859 |) A | Based | on | WO | 200069904 |
| ΕP | 1181315 | A1 | Based | on | WO | 200069904 |
| BR | 2000011294 | 1 A | Based | on | WO | 200069904 |

PRIORITY APPLN. INFO: US 1999-133973P 19990513

AN 2001-016210 [02] WPIDS

AB WO 200069904 A UPAB: 20010110

NOVELTY - Isolated or recombinant polypeptide (I) that comprises, mimics or cross-reacts with a B- or T-cell epitope of a FlgE (flagellar hook) polypeptide from a Lawsonia spp.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) a vaccine comprising, at least one carrier, diluent or adjuvant and a (I) that has at least 60% sequence identity overall with a fully defined 502 aa sequence (1), (given in the specification) or its immunogenic homolog, analog or derivative that is immunologically cross-reactive with L. intracellularis;
- (2) a vaccine vector comprising, in expressible form, a nucleic acid sequence (II) that encodes (1);
- (3) a poly- or mono-clonal antibody (Ab) that binds to Lawsonia FlgE polypeptide, or its derivatives, that have at least 60% sequence identity with (1);
- (4) an isolated nucleic acid (III) that encodes a peptide, oligopeptide or polypeptide having at least 60% sequence identity with (1) or its homolog, analog or derivative that mimics a B- or T-cell epitope, also complements of (III);
- (5) a probe or primer containing at least 15 contiguous nucleotides from a fully defined 1509 bp sequence (2), (given in the specification) or its complement; and
 - (6) a plasmid pALK11 (ATCC 207156).

ACTIVITY - Antibacterial.

MECHANISM OF ACTION - Induction of a specific humoral immune response. No data given.

USE - (I) are used as antigens in vaccines to prevent

or treat infection by Lawsonia, in birds and animals, especially pigs, to raise specific antibodies (Ab) and to detect past or present infection. Ab are also useful in diagnosis, to detect L. intracellularis or immunologically cross-reactive species (claimed), also for identification of epitopes in FlgE. Vectors that contain nucleic acid (II) that encodes (I) are also useful in genetic vaccines, and fragments of (II) are useful as primers or probes for detecting L. intracellularis or related microorganisms, in hybridization or amplification assays.

Dwg.0/1

L7 ANSWER 10 OF 21 WPIDS (C) 2003 THOMSON DERWENT

ACCESSION NUMBER: 2001-031924 [04] WPIDS

DOC. NO. CPI: C2001-009790

TITLE: Isolated or recombinant polypeptide for treating

porcine and avian species against Lawsonia

intracellularis infection, comprises,

mimics or cross-reacts with the B or T cell epitope

of Lawsonia SodC polypeptide.

DERWENT CLASS: B04 D16

INVENTOR(S): ANKENBAUER, R G; HASSE, D; PANACCIO, M; ROSEY, E L;

WRIGHT, C

PATENT ASSIGNEE(S): (AGRI-N) AGRIC VICTORIA SERVICES PTY LTD; (PFIZ)

PFIZER PROD INC; (PIGR-N) PIG RES & DEV CORP;

(AUPO-N) AUSTRALIAN PORK LTD

COUNTRY COUNT: 93

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2000069903 A1 20001123 (200104)* EN 85

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC

MW NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT

RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA $_{\rm ZW}$

AU 2000043858 A 20001205 (200113)

EP 1177212 A1 20020206 (200218) EN

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI

BR 2000011292 A 20020226 (200223)

JP 2003501013 W 20030114 (200306) 89

APPLICATION DETAILS:

| PATENT NO KIND | APPLICATION | DATE |
|------------------|----------------|----------|
| WO 2000069903 A1 | WO 2000-AU436 | 20000511 |
| AU 2000043858 A | AU 2000-43858 | 20000511 |
| EP 1177212 A1. | EP 2000-924975 | 20000511 |
| | WO 2000-AU436 | 20000511 |
| BR 2000011292 A | BR 2000-11292 | 20000511 |
| | WO 2000-AU436 | 20000511 |
| JP 2003501013 W | JP 2000-618319 | 20000511 |
| | WO 2000-AU436 | 20000511 |

FILING DETAILS:

| PAT | ENT NO K | IND | | | PAT | ENT NO |
|-----|------------|------------|-------|----|-----|-----------|
| AU | 2000043858 | А | Based | on | WO | 200069903 |
| EP | 1177212 | A 1 | Based | on | WO | 200069903 |
| ΒR | 2000011292 | Α | Based | on | WO | 200069903 |
| JΡ | 2003501013 | W | Based | on | WO | 200069903 |

PRIORITY APPLN. INFO: US 1999-133989P 19990513

AN 2001-031924 [04] WPIDS

AB WO 200069903 A UPAB: 20010118

NOVELTY - An isolated or recombinant immunogenic polypeptide (I) which comprises, mimics or cross-reacts with a B-cell or T-cell epitope of a Lawsonia SodC polypeptide, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) a vaccine composition (II) for the prophylaxis or treatment of infection of an animal by Lawsonia comprising an immunogenic component which comprises (I), which is immunologically cross-reactive with Lawsonia intracellularis and one or more carriers, diluents or adjuvants suitable for veterinary or pharmaceutical use;
- (2) a combination vaccine composition (III) for the prophylaxis or treatment of infection of an animal by Lawsonia comprising, a first immunogenic component which comprises (I), a second immunogenic component comprising an antigenic L. intracellularis peptide, polypeptide or protein and one or more carriers, diluents or adjuvants suitable for veterinary or pharmaceutical use;
- (3) a vaccine vector (IV) comprising, in an expressible form, an isolated nucleic acid molecule having a nucleotide sequence that encodes an isolated or recombinant immunogenic polypeptide which comprises the sequence (S) such that the immunogenic polypeptide is expressible at a level sufficient to confer immunity against Lawsonia, when administered to a porcine or avian animal;
- (4) a polyclonal or monoclonal antibody molecule (V) that is capable of binding specifically to (I);
- (5) an isolated nucleic acid molecule (VI) that encodes (I), or its complement;
- (6) a probe or primer (VII) having at least 15 contiguous nucleotides in length derived from the fully defined sequence of 543 base pairs (bp) as given in the specification or its complement; and

(7) a plasmid designated pALK14 (ATCC 207155).

ACTIVITY - Antibacterial.

No biological data is given.

MECHANISM OF ACTION - Vaccine.

No biological data is given.

USE - (I) is useful for diagnosing infection of a porcine or avian animal or identifying whether or not the animal has suffered from a past infection or is currently infected with L. intracellularis or a microorganism that is immunologically cross-reactive to it, by contacting whole serum, blood lymph nodes, ileum, caecum, small intestine, large intestine, feces or rectal swab derived from the animal with (V) or (I) for a time and under conditions sufficient for an antigen:antibody complex to form and detecting the complex formed. (VII) is useful for detecting

L. intracellularis or related microorganisms in a sample derived from the animal by hybridizing (VII) or its complement to the sample and then detecting the hybridization using a nucleic acid based hybridization or amplification reaction. (I) is useful in the preparation of a medicament for the treatment and prophylaxis of porcine proliferative enteropathy (PPE) in animals, particularly porcine or avian animals. (IV) is useful for producing a proteinaceous immunogenic component of (II) or (III) or is useful in a DNA vaccine. (II) and (III) are useful for treatment and/or prophylaxis of porcine and/or avian species against any bacterium belonging to the same serovar or serogroup as L.

Dwg.0/0

L7 ANSWER 11 OF 21 MEDLINE DUPLICATE 4

ACCESSION NUMBER: 2001041976 MEDLINE

DOCUMENT NUMBER: 20399396 PubMed ID: 10945299

TITLE: Immunohistochemistry and polymerase chain reaction

for the detection of Lawsonia

intracellularis in porcine intestinal tissues

with proliferative enteropathy.

AUTHOR: Kim J; Choi C; Cho W S; Chae C

CORPORATE SOURCE: Department of Veterinary Pathology, College of

Veterinary Medicine and School of Agricultural Biotechnology, Seoul National University, Suwon,

Kyounggi-Do, Republic of Korea.

SOURCE: JOURNAL OF VETERINARY MEDICAL SCIENCE, (2000 Jul) 62

(7) 771-3.

Journal code: 9105360. ISSN: 0916-7250.

PUB. COUNTRY: Japan

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200012

ENTRY DATE: Entered STN: 20010322

Last Updated on STN: 20010322

Entered Medline: 20001207

AB Detection method of Lawsonia intracellularis was studied in formalin-fixed paraffin-embedded intestinal tissues from 5 naturally infected pigs by immunohistochemistry with a monoclonal

antibody against outer membrane protein of L. intracellularis. Warthin-Starry silver

stain revealed clusters of argyrophilic, slightly curved rod-shaped organisms in the apical cytoplasm of enterocytes.

Immunohistochemical staining with a L.

intracellularis-specific monoclonal antibody confirmed the presence of the organism in the apical cytoplasm of hyperplastic enterocytes. The presence of L. intracellularis

in the ileum of pig with proliferative enteropathy was confirmed by polymerase chain reaction (PCR) further on the basis of amplification of 319 base pair products specific for porcine

L. intracellularis chromosomal DNA.

Immunohistochemistry and PCR may be a complementary method to confirm the diagnosis of L. intracellularis infection in pigs.

L7 ANSWER 12 OF 21 MEDLINE

ACCESSION NUMBER: 2000150299 MEDLINE

Searcher: Shears 308-4994

DUPLICATE 5

PubMed ID: 10684754 DOCUMENT NUMBER: 20150299

TITLE: Detection of Lawsonia

intracellularis in the tonsils of pigs with

proliferative enteropathy.

Jensen T K; Moller K; Lindecrona R; Jorsal S E AUTHOR: Danish Veterinary Laboratory, Copenhagen, Denmark. CORPORATE SOURCE: RESEARCH IN VETERINARY SCIENCE, (2000 Feb) 68 (1) SOURCE:

23-6.

Journal code: 0401300. ISSN: 0034-5288.

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

English LANGUAGE:

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200003

ENTRY DATE: Entered STN: 20000407

> Last Updated on STN: 20000407 Entered Medline: 20000329

AB The presence of Lawsonia intracellularis, the obligate intracellular bacterium causing proliferative enteropathy (PE), in the tonsils of pigs as a locus for infection or extraintestinal occurrence of the bacterium was investigated by PCR and immunohistochemistry. Tonsillar occurrence of L. intracellularis could be part of the pathogenesis of PE and an important risk factor in the spread of the disease. L. intracellularis was detected by only PCR in the tonsils of 2/32 pigs without PE at necropsy but with a clinical history of diarrhoea and detection of the bacterium in faeces 1 to 3 weeks prior to necropsy but not in four pigs with moderate PE lesions. However, L. intracellularis was detected in the tonsils of 4/9 pigs with PE complicated with necroses and in 4/4 pigs with proliferative haemorrhagic enteropathy in which L . intracellularis antigen also was demonstrated in tonsillar macrophages and as intact bacteria in the lumen of the crypts. The results show that L. intracellularis is detectable in the tonsils of pigs and that the tonsillar presence of L. intracellularis appears to be correlated to the severity of the intestinal lesions possibly as a result of

local retention and not as part of the pathogenesis of PE. Copyright 2000 Harcourt Publishers LtdCopyright 2000 Harcourt Publishers Ltd.

ANSWER 13 OF 21 SCISEARCH COPYRIGHT 2003 ISI (R)

ACCESSION NUMBER: 1999:172294 SCISEARCH

THE GENUINE ARTICLE: 169GX

Attempted infection of mice, rats and chickens by TITLE:

porcine strains of Lawsonia

intracellularis

Collins A M (Reprint); Love R J; Jasni S; McOrist S AUTHOR:

CORPORATE SOURCE: UNIV SYDNEY, DEPT VET CLIN SCI, CAMDEN, NSW 2570, AUSTRALIA (Reprint); UNIV EDINBURGH, DEPT VET

PATHOL, EDINBURGH EH8 9YL, MIDLOTHIAN, SCOTLAND; VPS

VETLAB, GLENSIDE, SA 5065, AUSTRALIA

COUNTRY OF AUTHOR:

AUSTRALIA; SCOTLAND

AUSTRALIAN VETERINARY JOURNAL, (FEB 1999) Vol. 77, SOURCE:

No. 2, pp. 120-122.

Publisher: AUSTRALIAN VETERINARY ASSN, 272 BRUNSWICK

RD BRUNSWICK, MELBOURNE VIC 3056, AUSTRALIA.

ISSN: 0005-0423.

Searcher : 308-4994 Shears

DOCUMENT TYPE: Article; Journal

FILE SEGMENT: AGRI LANGUAGE: English

REFERENCE COUNT: 17

•

L7 ANSWER 14 OF 21 MEDLINE

ACCESSION NUMBER: 1998281713 MEDLINE

DOCUMENT NUMBER: 98281713 PubMed ID: 9620403

TITLE: Proliferative enterocolitis associated with dual

infection with enteropathogenic Escherichia coli and

Lawsonia intracellularis in

rabbits.

AUTHOR: Schauer D B; McCathey S N; Daft B M; Jha S S;

Tatterson L E; Taylor N S; Fox J G

CORPORATE SOURCE: Division of Comparative Medicine, Massachusetts

Institute of Technology, Cambridge 02139, USA..

schauer@mit.edu

CONTRACT NUMBER: CA63112 (NCI)

RR07036 (NCRR)

SOURCE: JOURNAL OF CLINICAL MICROBIOLOGY, (1998 Jun) 36 (6)

1700-3.

Journal code: 7505564. ISSN: 0095-1137.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199807

ENTRY DATE: Entered STN: 19980811

Last Updated on STN: 19980811 Entered Medline: 19980728

AB Both enteropathogenic Escherichia coli (EPEC) and an obligate intracellular bacterium, previously referred to as an intracellular

Campylobacter-like organism and now designated Lawsonia

intracellularis, have been reported as causes of
enterocolitis in rabbits. An outbreak of enterocolitis in a group of
rabbits, characterized by an unusually high rate of mortality, was
found to be associated with dual infection with EPEC and L
. intracellularis. The EPEC strain was found to have eaeA
gene homology but was negative for afrA homology. The absence of the
afrA gene, which encodes the structural subunit for the AF/R1 pilus,
indicates that this rabbit EPEC strain is distinct from the
prototypic RDEC-1 strain. This finding suggests that rabbit EPEC
strains widely reported in Western Europe, which lack AF/R1 pili,
are also present in rabbits in the United States. Dual infection
with these two pathogens in rabbits has not been previously reported
and may have contributed to the unusually high mortality observed in
this outbreak.

L7 ANSWER 15 OF 21 MEDLINE DUPLICATE 6

ACCESSION NUMBER: 1998348117 MEDLINE

DOCUMENT NUMBER: 98348117 PubMed ID: 9684975

TITLE: Subclinical proliferative enteropathy in sentinel

rabbits associated with Lawsonia

intracellularis.

AUTHOR: Duhamel G E; Klein E C; Elder R O; Gebhart C J

CORPORATE SOURCE: Department of Veterinary and Biomedical Sciences,

University of Nebraska, Lincoln 68583-0905, USA.

SOURCE: VETERINARY PATHOLOGY, (1998 Jul) 35 (4) 300-3.

Journal code: 0312020. ISSN: 0300-9858.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199809

ENTRY DATE: Entered STN: 19981008

Last Updated on STN: 19981008 Entered Medline: 19980930

AB Light microscopic and ultrastructural changes of naturally acquired proliferative enteropathy were observed in two of three young sentinel New Zealand White rabbits. The etiologic agent, Lawsonia intracellularis, was demonstrated in the tissues using morphologic, immunohistochemical, and molecular methods. Proliferative enteropathy was associated with infection of villous and crypt enterocytes by intracellular organisms genotypically and antigenically related to L. intracellularis of various other animal species.

L7 ANSWER 16 OF 21 VETU COPYRIGHT 2003 THOMSON DERWENT

ACCESSION NUMBER: 1998-63015 VETU

TITLE: Control of infectious enteric diseases of swine.

AUTHOR: Lanza I CORPORATE SOURCE: Elanco

LOCATION: Madrid, Esp.

SOURCE: Proc.Int.Pig Vet.Soc.Congress (15 Meet., Pt. 1, 79-85,

1998) 1 Fig. 45 Ref.

AVAIL. OF DOC.: Elanco Sanidad Animal, Madrid, Spain.

LANGUAGE: English
DOCUMENT TYPE: Journal
FIELD AVAIL.: AB; LA; CT

AN 1998-63015 VETU

The control of infectious enteric diseases in pigs is reviewed, with respect to transmissible gastroenteritis (TGE), porcine epidemic diarrhea (PED), rotavirus infection, colibacillosis (caused by enterotoxigenic E. coli), coccidiosis (Isospora suis), salmonellosis (Salm. typhimurium or derby), Clostr. perfringens infection, swine dysentery (Serpulina hyodysenteriae), colonic spirochaetosis (S. pilosicoli), porcine proliferative enteropathy (Lawsonia intracellularis). Control involves the pig herd (maintain free of disease by controlling new stock), farm management practices (good hygiene, pig flow), the vaccination program and strategic medications. The aim is to reduce the incidence of the disease to the lowest levels possible, thus making the use of therapeutic medication to cure sick animals rare. (conference paper).

ABEX If TGE strikes neonates can be protected by raising the immune status of the sow by exposure to virus and production of antibodies in colostrum and milk. PED experimental attenuated vaccines show promise. Rotavirus is usually endemic; environmental virus levels should be kept low and passive transfer of immunity from the dam should be maximized by vaccines. Vaccination of pregnant sows with appropriate serotype vaccines will protect the neonate against E. coli. Postweaning, edema disease can be controlled by verotoxin toxoids but parenteral fimbrial antigen vaccines are not effective. Probiotics, antibiotics (apramycin, colistin, neomycin), organic acids or zinc oxide can be added to weaner feed. Anticoccidial therapy of piglets before diarrhea occurs helps

control coccidiosis. Supportive rehydration and mass medication helps against Salm.; killed vaccines are of little use; continuous in-feed antibiotics are not recommended. C. perfringens type C necrotic enteritis is controlled by vaccination of pregnant sows; antitoxin can be helpful. Vaccines do not contain toxins against type A enteritis; colostrum is usually effective in neonates, and in weaned pigs an antibiotic in feed is effective. S. hyodysenteriae shows resistance to dimetridazole and lincomycin; carbadox, tiamulin and pleuromulins are better. Bivalent inactivated vaccines reduce symptoms and induce a lactogenic immunity. L. intracellularis cannot be eradicated but in feed tylosin, tiamulin or chlortetracycline can control it.

L7 ANSWER 17 OF 21 MEDLINE DUPLICATE 7

ACCESSION NUMBER: 97370587 MEDLINE

DOCUMENT NUMBER: 97370587 PubMed ID: 9226893

TITLE: Comparison of the 16S ribosomal DNA sequences from

the intracellular agents of proliferative enteritis in a hamster, deer, and ostrich with the sequence of

a porcine isolate of Lawsonia

intracellularis.

AUTHOR: Cooper D M; Swanson D L; Barns S M; Gebhart C J

CORPORATE SOURCE: Division of Comparative Medicine, Research Animal

Resources, Medical School, University of Minnesota,

Minneapolis 55455, USA.

SOURCE: INTERNATIONAL JOURNAL OF SYSTEMATIC BACTERIOLOGY,

(1997 Jul) 47 (3) 635-9.

Journal code: 0042143. ISSN: 0020-7713.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

OTHER SOURCE: GENBANK-U65995; GENBANK-U65996; GENBANK-U65997;

GENBANK-U65998

ENTRY MONTH: 199708

ENTRY DATE: Entered STN: 19970908

Last Updated on STN: 19990129 Entered Medline: 19970826

AB Proliferative enteritis is an enteric disease that affects a variety of animals. The causative agent in swine has been determined to be an obligate intracellular bacterium, Lawsonia

intracellularis, related to the sulfate-reducing bacterium
Desulfovibrio desulfuricans. The intracellular agents found in the
lesions of different animal species are antigenically
similar. In addition, strains from the pig, ferret, and hamster have
been shown to be genetically similar. In this study we performed a
partial 16S ribosomal DNA sequence analysis on the intracellular.
agent of proliferative enteritis from a hamster, a deer, and an
ostrich and compared these sequences to that of the porcine
L. intracellularis isolate. Results of this study

indicate that the intracellular agents from these species with proliferative enteritis have high sequence similarity, indicating that they are all in the genus Lawsonia and that they may also be the same species, L. intracellularis.

the same species, I. Intlatellatella

ACCESSION NUMBER: 97254956 MEDLINE

ANSWER 18 OF 21

Searcher : Shears 308-4994

MEDLINE

DOCUMENT NUMBER: 97254956 PubMed ID: 9100338

TITLE: In-vitro interactions of Lawsonia

intracellularis with cultured enterocytes. McOrist S; Mackie R A; Lawson G H; Smith D G AUTHOR: Department of Veterinary Pathology, University of CORPORATE SOURCE:

Edinburgh, Easter Bush, Midlothian, UK.

VETERINARY MICROBIOLOGY, (1997 Mar) 54 (3-4) 385-92. SOURCE:

Journal code: 7705469. ISSN: 0378-1135.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

English LANGUAGE:

Priority Journals FILE SEGMENT:

199706 ENTRY MONTH:

Entered STN: 19970630 ENTRY DATE:

> Last Updated on STN: 20000303 Entered Medline: 19970619

Strains of the obligately intracellular bacterium Lawsonia AΒ

intracellularis, the etiologic agent of porcine proliferative enteropathy, were co-cultured in rat enterocyte cell cultures (IEC-18) and examined ultrastructurally. No regular surface arrays typical of surface or S-layers were visible on any bacterial strain, with or without Triton-X-100 detergent treatment. In

separate experiments, there was no difference in the ability of L. intracellularis to attach and enter enterocytes

with or without the presence of added bovine plasma fibronectin, or the peptide Arg-Gly-Ser. Interestingly, there was an increase in the invasiveness of L. intracellularis in the

presence of the peptide Arg-Gly-Asp (RGD), in a dose-related manner.

A reduction was observed in the ability of L.

intracellularis to invade enterocytes in the presence of monovalent fragments of IgG monoclonal antibodies to an outer surface component of L. intracellularis. This neutralization showed an antibody concentration-dependent titration

effect and was not apparent with co-cultures incorporating control antibodies. The exact nature of ligand and cell receptor interactions for L. intracellularis remain to be

determined.

AUTHOR:

DUPLICATE 8 ANSWER 19 OF 21 MEDLINE 1.7

ACCESSION NUMBER: 97218646 MEDLINE

PubMed ID: 9066083 DOCUMENT NUMBER: 97218646

Intracellular Campylobacter-like organisms associated TITLE:

with rectal prolapse and proliferative

enteroproctitis in emus (Dromaius novaehollandiae). Lemarchand T X; Tully T N Jr; Shane S M; Duncan D E

Department of Pathology, School of Veterinary CORPORATE SOURCE:

Medicine, Louisiana State University, Baton Rouge

70803, USA.

VETERINARY PATHOLOGY, (1997 Mar) 34 (2) 152-6. SOURCE:

Journal code: 0312020. ISSN: 0300-9858.

United States PUB. COUNTRY:

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

Priority Journals FILE SEGMENT:

199705 ENTRY MONTH:

Entered STN: 19970602 ENTRY DATE:

Last Updated on STN: 20000303 Entered Medline: 19970522

Rectal prolapse was the presenting clinical finding in a group of AB juvenile emus (Dromaius novaehollandiae). Gross findings included severely thickened and rugose distal rectal mucosae. Histologically, there were thickened villi, enterocyte hyperplasia, dilated glands filled with mucus and heterophils, and a dense infiltrate of heterophils, macrophages, lymphocytes, and plasma cells in the lamina propria. Examination of Warthin-Starry silver-stained sections revealed numerous apically located comma-shaped intracytoplasmic bacteria approximately 1 x 3 microns in size. Campylobacter-like organisms morphologically compatible with ileal symbiont intracellularis now known as Lawsonia intracellularis were seen via electron microscopy. Bacteria were further characterized by indirect immunofluorescence using monoclonal antibody specific for the 25-27-kd outer membrane protein of L. intracellularis.

L7 ANSWER 20 OF 21 CABA COPYRIGHT 2003 CABI

ACCESSION NUMBER: 96:105329 CABA

DOCUMENT NUMBER: 962209126

TITLE: The pathogenesis of necrotic proliferative

colitis in swine is linked to whipworm induced

suppression of mucosal immunity to resident

bacteria

AUTHOR: Mansfield, L. S.; Urban, J. F., Jr.

CORPORATE SOURCE: College of Veterinary Medicine, Michigan State

University, East Lansing, MI 48824, USA. Veterinary Immunology and Immunopathology, (1996) Vol. 50, No. 1/2, pp. 1-17. 30 ref.

ISSN: 0165-2427

DOCUMENT TYPE: Journal LANGUAGE: English

SOURCE:

The pathogenesis of mucohaemorrhagic enteritis syndrome was AB investigated using 4 groups of pigs. Group 1 were inoculated with 2500 embryonated Trichuris suis eggs alone, while group 2 received T. suis eggs along with broad spectrum antibiotic treatment. Two control groups were not inoculated and were either treated with antibiotic or untreated. Group 1 pigs exhibited diarrhoea, mucosal oedema, inflammatory cell infiltration, bacterial accumulation at the site of worm attachment in the proximal colon, and intestinal adenomatosis associated with the intracellular Ileal symbiont intracellularis [Lawsonia intracellularis] bacteria. In addition, enlarged lymphoglandular complexes (LGCs) containing numerous extracellular bacteria, eosinophils, lymphocytes, macrophages and neutrophils were observed in the distal colon. Group 2 pigs had lesions localized to the site of worm attachment and histologically normal LGCs with no invasive bacteria in the distal colon. The control pigs, with or without antibiotic treatment, exhibited no pathology or bacterial invasion. It is concluded that the complex pathogenesis of necrotic proliferative colitis in pigs may be linked to worm induced suppression of mucosal immunity to resident bacteria. The association between bacteria, lymphocytes and macrophages in the LGCs of group 1 pigs suggests an antigen-processing role for these structures in the colon.

L7 ANSWER 21 OF 21 MEDLINE DUPLICATE 9

ACCESSION NUMBER: 85081914 MEDI:INE

DOCUMENT NUMBER: 85081914 PubMed ID: 6096477

TITLE: Inherited deficiency of the Mac-1, LFA-1, p150,95

glycoprotein family and its molecular basis.

AUTHOR: Springer T A; Thompson W S; Miller L J; Schmalstieg F

C; Anderson D C

CONTRACT NUMBER: AI 19031 (NIAID)

CA 31798 (NCI) CA 31799 (NCI)

SOURCE: JOURNAL OF EXPERIMENTAL MEDICINE, (1984 Dec 1) 160

(6) 1901-18.

Journal code: 2985109R. ISSN: 0022-1007.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198501

L8

L9

ENTRY DATE: Entered STN: 19900320

Last Updated on STN: 19970203 Entered Medline: 19850128

Leukocyte surface glycoproteins that share a common beta subunit ΑB have been found to be congenitally deficient in three unrelated patients with recurring bacterial infection. The glycoproteins, Mac-1, LFA-1, and p150,95, have the subunit compositions alpha M beta, alpha L beta, and alpha X beta, respectively. Using subunit-specific monoclonal antibodies, both the alpha M and beta subunits of Mac-1, the alpha L and beta subunits of LFA-1, and at the least the beta subunit of p150,95, were found to be deficient at the cell surface by the techniques of immunofluorescence flow cytometry, radioimmunoassay, and immunoprecipitation. A latent pool of Mac-1 that can be expressed on granulocyte surfaces in response to secretory stimuli, such as f-Met-Leu-Phe, was also lacking in patients. Deficiency was found on all leukocytes tested, including granulocytes, monocytes, and T and B lymphocytes. Quantitation by immunofluorescence cytometry of subunits on granulocytes from parents of these patients and of a fourth deceased patient showed approximately half-normal surface expression, and, together with data on other siblings and a family with an affected father and children, demonstrate autosomal recessive inheritance. Deficiency appears to be quantitative rather than qualitative, with two patients expressing approximately 0.5% and one patient approximately 5% of normal amounts. The latter patient had alpha beta complexes on the cell surface detectable by immunoprecipitation. Biosynthesis experiments showed the presence of normal amounts of alpha'L intracellular precursor in lymphoid lines of all three patients. Together with surface deficiency of three molecules that share a common beta subunit but have differing alpha subunits, this suggests the primary deficiency is of the beta subunit. The lack of maturation of alpha'L to alpha L and the deficiency of the alpha subunits at the cell surface and in latent pools suggests that association with the beta subunit is required for alpha subunit processing and transport to the cell surface or to latent pools. The molecular basis of this disease is discussed in light of adhesion-related functional abnormalities in patients' leukocytes and the blockade of similar functions in healthy cells by monoclonal antibodies.

(FILE 'MEDLINE' ENTERED AT 11:23:21 ON 10 APR 2003)

15 SEA FILE=MEDLINE ABB=ON PLU=ON LAWSONIA/CT

48331 SEA FILE=MEDLINE ABB=ON PLU=ON ANTIGENS/CT

| L10 0 SEA FILE=MEDLINE ABB=ON PLU=ON L8 AND L9 |
|--|
| L8 15 SEA FILE=MEDLINE ABB=ON PLU=ON LAWSONIA/CT L11 9492 SEA FILE=MEDLINE ABB=ON PLU=ON "BACTERIAL OUTER MEMBRANE PROTEINS"/CT L12 0 SEA FILE=MEDLINE ABB=ON PLU=ON L8 AND L11 |
| LIZ . 0 SEA FILE-MEDDINE ADD-ON FRO-ON RO AND DIT |
| L8 15 SEA FILE=MEDLINE ABB=ON PLU=ON LAWSONIA/CT L13 6043 SEA FILE=MEDLINE ABB=ON PLU=ON VACCINES/CT L14 0 SEA FILE=MEDLINE ABB=ON PLU=ON L8 AND L13 |
| FILE 'USPATFULL' ENTERED AT 11:26:03 ON 10 APR 2003 L15 13 SEA ABB=ON PLU=ON ((LAWSON? OR L)(W)INTRACELL?)(L)(OMP OR OUTER MEMBRAN? PROTEIN OR ANTIGEN##) |
| L15 ANSWER 1 OF 13 USPATFULL ACCESSION NUMBER: 2003:29860 USPATFULL TITLE: Lawsonia intracellularis proteins, and related methods and materials |
| INVENTOR(S): Rosey, Everett L., Preston, CT, UNITED STATES |
| NUMBER KIND DATE |
| NUMBER DATE |
| NUMBER DATE |
| PRIORITY INFORMATION: US 1999-160922P 19991022 (60) US 1999-163858P 19991105 (60) DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION LEGAL REPRESENTATIVE: KOHN & ASSOCIATES, PLLC, SUITE 410, 30500 |
| PRIORITY INFORMATION: US 1999-160922P 19991022 (60) US 1999-163858P 19991105 (60) DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION |
| PRIORITY INFORMATION: US 1999-160922P 19991022 (60) US 1999-163858P 19991105 (60) DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION LEGAL REPRESENTATIVE: KOHN & ASSOCIATES, PLLC, SUITE 410, 30500 NORTHWESTERN HWY., FARMINGTON HILLS, MI, 48334 NUMBER OF CLAIMS: 20 EXEMPLARY CLAIM: 1 NUMBER OF DRAWINGS: 8 Drawing Page(s) |
| PRIORITY INFORMATION: US 1999-160922P 19991022 (60) US 1999-163858P 19991105 (60) DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION LEGAL REPRESENTATIVE: KOHN & ASSOCIATES, PLLC, SUITE 410, 30500 NORTHWESTERN HWY., FARMINGTON HILLS, MI, 48334 NUMBER OF CLAIMS: 20 EXEMPLARY CLAIM: 1 NUMBER OF DRAWINGS: 8 Drawing Page(s) LINE COUNT: 3947 CAS INDEXING IS AVAILABLE FOR THIS PATENT. AB Isolated polynucleotide molecules contain a nucleotide sequence that encodes a L. intracellularis HtrA, PonA, HypC, LysS, YcfW, ABC1, or Omp100 protein, a substantial portion of the sequences, or a homologous sequence. Related polypeptides, immunogenic compositions and assays are described. CAS INDEXING IS AVAILABLE FOR THIS PATENT. |
| PRIORITY INFORMATION: US 1999-160922P 19991022 (60) US 1999-163858P 19991105 (60) DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION LEGAL REPRESENTATIVE: KOHN & ASSOCIATES, PLLC, SUITE 410, 30500 NORTHWESTERN HWY., FARMINGTON HILLS, MI, 48334 NUMBER OF CLAIMS: 20 EXEMPLARY CLAIM: 1 NUMBER OF DRAWINGS: 8 Drawing Page(s) LINE COUNT: 3947 CAS INDEXING IS AVAILABLE FOR THIS PATENT. AB Isolated polynucleotide molecules contain a nucleotide sequence that encodes a L. intracellularis HtrA, PonA, HypC, LysS, YcfW, ABC1, or Omp100 protein, a substantial portion of the sequences, or a homologous sequence. Related polypeptides, immunogenic compositions and assays are described. CAS INDEXING IS AVAILABLE FOR THIS PATENT. INCL INCLM: 424/190.100 INCLS: 435/219.000; 435/320.100; 435/252.300; 536/023.200; |
| PRIORITY INFORMATION: US 1999-160922P 19991022 (60) US 1999-163858P 19991105 (60) DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION LEGAL REPRESENTATIVE: KOHN & ASSOCIATES, PLLC, SUITE 410, 30500 NORTHWESTERN HWY., FARMINGTON HILLS, MI, 48334 NUMBER OF CLAIMS: 20 EXEMPLARY CLAIM: 1 NUMBER OF DRAWINGS: 8 Drawing Page(s) LINE COUNT: 3947 CAS INDEXING IS AVAILABLE FOR THIS PATENT. AB Isolated polynucleotide molecules contain a nucleotide sequence that encodes a L. intracellularis HtrA, PonA, HypC, LysS, YcfW, ABC1, or Omp100 protein, a substantial portion of the sequences, or a homologous sequence. Related polypeptides, immunogenic compositions and assays are described. CAS INDEXING IS AVAILABLE FOR THIS PATENT. INCL INCLM: 424/190.100 |

ACCESSION NUMBER: 2002:251933 USPATFULL

Protein scaffold and its use to multimerise TITLE:

monomeric polypeptides

INVENTOR(S): Hill, Fergal Conan, Les Martres de Veyre, FRANCE

Chatellier, Jean, Les Martres de Veyre, FRANCE Fersht, Alan Roy, Cambridge, UNITED KINGDOM

NUMBER KIND DATE

US 2002137891 A1 20020926 US 2001-7628 A1 20011108 (10) PATENT INFORMATION: APPLICATION INFO.:

Continuation-in-part of Ser. No. WO 2000-GB1815, RELATED APPLN. INFO.:

filed on 5 Dec 2000, UNKNOWN

DATE NUMBER -----GB 1999-11298 19990514 PRIORITY INFORMATION: GB 1999-28788 19991203

GB 1999-28831 19991206 Utility DOCUMENT TYPE:

FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: FROMMER LAWRENCE & HAUG, 745 FIFTH AVENUE- 10TH

FL., NEW YORK, NY, 10151

NUMBER OF CLAIMS: 36 EXEMPLARY CLAIM: 1

12 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT: 2025

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to polypeptide monomer capable of

oligomerisation, said monomer comprising a heterologous amino acid

sequence inserted into the sequence of a subunit of an

oligomerisable protein scaffold.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCLM: 530/350.000

INCLS: 514/012.000; 514/013.000; 514/014.000; 514/015.000;

514/021.000; 536/023.400

NCL NCLM: 530/350.000

NCLS: 514/012.000; 514/013.000; 514/014.000; 514/015.000;

514/021.000; 536/023.400

L15 ANSWER 3 OF 13 USPATFULL

2002:136784 USPATFULL ACCESSION NUMBER:

TITLE: Staphylococcus aureus genes and polypeptides INVENTOR(S): Bailey, Camella, Washington, DC, United States

Choi, Gil H., Rockville, MD, United States

Human Genome Sciences, Inc., Rockville, MD, PATENT ASSIGNEE(S):

United States (U.S. corporation)

NUMBER KIND DATE US 6403337 US 2000-512255 В1 20020611 PATENT INFORMATION: 20000224 (9) APPLICATION INFO.:

Continuation-in-part of Ser. No. WO 1999-US19726, RELATED APPLN. INFO.:

filed on 31 Aug 1999 Continuation-in-part of Ser.

No. US 1997-956171, filed on 20 Oct 1997

Continuation-in-part of Ser. No. US 1997-781986, filed on 3 Jan 1997 Continuation-in-part of Ser.

No. US 1997-781986, filed on 5 Jan 1997

Continuation-in-part of Ser. No. US 1997-781986,

filed on 5 Jan 1997

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Brusca, John S.

LEGAL REPRESENTATIVE: Human Genome Sciences, Inc.

NUMBER OF CLAIMS: 65 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 0 Drawing Figure(s); 0 Drawing Page(s)

LINE COUNT: 6784

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to novel genes from S. aureus and the polypeptides they encode. Also provided as are vectors, host cells, antibodies and recombinant methods for producing the same. The invention further relates to screening methods for identifying agonists and antagonists of S. aureus polypeptide activity. The invention additionally relates to diagnostic methods for detecting Staphylococcus nucleic acids, polypeptides and antibodies in a biological sample. The present invention further relates to novel vaccines for the prevention or attenuation of infection by Staphylococcus.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCL INCLM: 435/069.700

INCLS: 435/468.000; 435/252.300; 435/320.100; 536/023.700

NCL NCLM: 435/069.700

NCLS: 435/252.300; 435/320.100; 435/468.000; 536/023.700

L15 ANSWER 4 OF 13 USPATFULL

ACCESSION NUMBER: 2002:61250 USPATFULL

TITLE: Methods of treating hepatitis delta virus

infection with beta-1-2'-deoxy-nucleosides

INVENTOR(S): Sommadossi, Jean-Pierre, Birmingham, AL, UNITED

STATES

Bryant, Martin L., Carlisle, MA, UNITED STATES

NUMBER DATE

PRIORITY INFORMATION: US 2000-207538P 20000526 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: KING & SPALDING, 191 PEACHTREE STREET, N.E.,

ATLANTA, GA, 30303-1763

NUMBER OF CLAIMS: 23 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 5 Drawing Page(s)

LINE COUNT: 2315

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A method and composition for treating a host infected with hepatitis D comprising administering an effective hepatitis D treatment amount of a described 2'-deoxy-.beta.-L-erythropentofuranonucleoside or a pharmaceutically acceptable salt or

prodrug thereof.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCLM: 514/045.000 INCL

INCLS: 514/046.000; 514/050.000; 514/047.000; 514/048.000;

514/051.000

NCL NCLM: 514/045.000

NCLS: 514/046.000; 514/050.000; 514/047.000; 514/048.000;

514/051.000

L15 ANSWER 5 OF 13 USPATFULL

2000:149713 USPATFULL ACCESSION NUMBER:

Methods for modulating T cell survival by TITLE:

modulating bcl-X.sub.L protein level

June, Carl H., 7 Harlow Ct., Rockville, MD, INVENTOR(S):

United States 20850

Thompson, Craig B., 1375 E. 57th St., Chicago,

IL, United States 60637

KIND NUMBER DATE ______

US 6143291 20001107 PATENT INFORMATION: US 1995-481739 19950607 (8) APPLICATION INFO.:

Continuation-in-part of Ser. No. US 1995-435518, RELATED APPLN. INFO.:

filed on 4 May 1995, now abandoned

DOCUMENT TYPE: Utility Granted FILE SEGMENT:

Hauda, Karen M. PRIMARY EXAMINER:

Lahive & Cockfield, LLP LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: 5 1,3 EXEMPLARY CLAIM:

21 Drawing Figure(s); 13 Drawing Page(s) NUMBER OF DRAWINGS:

2507 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Methods for protecting a T cell from cell death are described. The AB methods involve contacting the T cell with an agent which augments the bcl-X.sub.L protein level in the T cell such that it is protected from cell death. The invention further pertains to methods for increasing the susceptibility of a T cell to cell death, comprising contacting the T cell with at least one agent which decreases bcl-X.sub.L protein level in the T cell. Both in vivo and in vitro methods are described.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCLM: 424/093.210 INCL

INCLS: 435/375.000; 435/320.100; 435/172.300

NCLM: 424/093.210 NCL

NCLS: 435/320.100; 435/375.000; 435/455.000

L15 ANSWER 6 OF 13 USPATFULL

ACCESSION NUMBER: 1999:36943 USPATFULL

Lawsonia intracellularis cultivation, TITLE:

anti-Lawsonia intracellularis vaccines and

diagnostic agents

Knittel, Jeffrey P., Ames, IA, United States INVENTOR(S):

Roof, Michael B., Ames, IA, United States

NOBL Laboratories, Inc., Sioux Center, IA, United PATENT ASSIGNEE(S):

States (U.S. corporation)

308-4994 Searcher : Shears

NUMBER KIND DATE _____ US 5885823 US 1996-658194 19990323 PATENT INFORMATION: 19960604 (8) APPLICATION INFO.: Continuation-in-part of Ser. No. US 1995-465337, RELATED APPLN. INFO.: filed on 5 Jun 1995, now patented, Pat. No. US 5714375 Utility DOCUMENT TYPE: Granted FILE SEGMENT: Hutzell, Paula K. PRIMARY EXAMINER: Masood, Khalid ASSISTANT EXAMINER: LEGAL REPRESENTATIVE: Dickstein Shapiro Morin & Oshinsky LLP NUMBER OF CLAIMS: 17 EXEMPLARY CLAIM: 1 1540 LINE COUNT: CAS INDEXING IS AVAILABLE FOR THIS PATENT. A method for large scale cultivation and attenuation of L. intracellularis bacteria by inoculating cells with L. intracellularis bacteria to infect the cells, incubating the infected cells in a reduced oxygen concentration and maintaining the infected cells in suspension. Anti-L. intracellularis vaccines are prepared from cultures grown in suspension. Diagnostic agents are also disclosed. CAS INDEXING IS AVAILABLE FOR THIS PATENT. INCLM: 435/243.000 INCL INCLS: 435/245.000; 435/252.100; 435/366.000; 435/383.000; 435/395.000; 435/403.000; 424/093.400; 424/234.100; 424/825.000 NCLM: 435/243.000 NCL NCLS: 424/093.400; 424/234.100; 424/825.000; 435/245.000; 435/252.100; 435/366.000; 435/383.000; 435/395.000; 435/403.000 L15 ANSWER 7 OF 13 USPATFULL ACCESSION NUMBER: 1998:65267 USPATFULL Method of treating catabolic, gut-associated TITLE: pathological processes and impaired host defenses Smith, Robert J., Brookline, MA, United States Wilmore, Douglas, Brookline, MA, United States INVENTOR(S): Brigham and Women's Hospital, Boston, MA, United PATENT ASSIGNEE(S): States (U.S. corporation) NUMBER KIND ______ US 5763485 US 1995-402827 19980609 PATENT INFORMATION: APPLICATION INFO.: 19950313 (8) Division of Ser. No. US 1993-51941, filed on 26 RELATED APPLN. INFO.: Apr 1993, now patented, Pat. No. US 5397803, issued on 14 Mar 1995 which is a continuation of Ser. No. US 1993-845819, filed on 9 Mar 1993, now abandoned which is a continuation-in-part of Ser. No. US 1989-360839, filed on 2 Jun 1989, now abandoned which is a continuation-in-part of Ser. No. US 1986-906530, filed on 12 Sep 1986, now patented, Pat. No. US 4857555 which is a

Searcher: Shears 308-4994

continuation-in-part of Ser. No. US 1985-775214,

filed on 12 Sep 1985, now abandoned

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

Raymond, Richard L. PRIMARY EXAMINER:

Sterne, Kessler, Goldstein & Fox P.L.L.C. LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 12 Drawing Figure(s); 9 Drawing Page(s)

LINE COUNT: 4313

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to a method for treating catabolic, gut-associated pathological processes including intestinal mucosal and pancreatic atrophy and enhanced gut permeability, impairment of host defenses and compromised immune function, and for promoting recovery from bone marrow transplantation in an animal, which comprises administering to an animal a therapeutically effective amount of glutamine or an analogue thereof.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCL INCLM: 514/563.000 NCL NCLM: 514/563.000

L15 ANSWER 8 OF 13 USPATFULL

ACCESSION NUMBER: 97:101796 USPATFULL

TITLE: Method of treating pancreatic atrophy

Smith, Robert J., Brookline, MA, United States INVENTOR(S):

Wilmore, Douglas, Brookline, MA, United States

Brigham and Women's Hospital, Boston, MA, United PATENT ASSIGNEE(S):

States (U.S. corporation)

DATE NUMBER KIND _____ US 5684045 US 1996-643937 19971104 PATENT INFORMATION: APPLICATION INFO.: 19960507 (8)

Division of Ser. No. US 1995-402827, filed on 13 RELATED APPLN. INFO.:

Mar 1995 which is a division of Ser. No. US 1993-51941, filed on 26 Apr 1993, now patented,

Pat. No. US 5397803 which is a

continuation-in-part of Ser. No. US 1992-845819, filed on 9 Mar 1992, now abandoned which is a continuation-in-part of Ser. No. US 1989-360839, filed on 2 Jun 1989, now abandoned which is a continuation-in-part of Ser. No. US 1986-906530, filed on 12 Sep 1986, now patented, Pat. No. US 4857555 which is a continuation-in-part of Ser. No. US 1985-775214, filed on 12 Sep 1985, now

abandoned Utility

DOCUMENT TYPE: FILE SEGMENT: Granted

Raymond, Richard L. PRIMARY EXAMINER:

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 12 Drawing Figure(s); 9 Drawing Page(s)

LINE COUNT: 4178

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to a method for treating catabolic,

gut-associated pathological processes including intestinal mucosal and pancreatic atrophy and enhanced gut permeability, impairment

> 308-4994 Searcher : Shears

of host defenses and compromised immune function, and for promoting recovery from bone marrow transplantation in an animal, which comprises administering to an animal a therapeutically effective amount of glutamine or an analogue thereof.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCLM: 514/563.000 NCL NCLM: 514/563.000

L15 ANSWER 9 OF 13 USPATFULL

97:18200 USPATFULL ACCESSION NUMBER:

Method of treating catabolic, gut-associated TITLE:

pathological processes and impaired host defenses

Smith, Robert J., Brookline, MA, United States INVENTOR(S):

Wilmore, Douglas, Brookline, MA, United States

Brigham and Women's Hospital, Boston, MA, United PATENT ASSIGNEE(S):

States (U.S. corporation)

KIND DATE NUMBER ______

US 5607975 US 5607975 19970304 US 1996-643939 19960507 (8) PATENT INFORMATION: APPLICATION INFO.:

Division of Ser. No. US 1995-402827, filed on 13 RELATED APPLN. INFO.:

> Mar 1995 which is a division of Ser. No. US 1993-51941, filed on 26 Apr 1993, now patented,

Pat. No. US 5397803 which is a

continuation-in-part of Ser. No. US 1992-845819, filed on 9 Mar 1992, now abandoned which is a continuation-in-part of Ser. No. US 1989-360839, filed on 2 Jun 1989, now abandoned which is a continuation-in-part of Ser. No. US 1986-906530, filed on 12 Sep 1986, now patented, Pat. No. US 4857555 which is a continuation-in-part of Ser. No. US 1985-775214, filed on 12 Sep 1985, now

abandoned

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Raymond, Richard L.

Sterne, Kessler, Goldstein & Fox P.L.L.C. LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 12 Drawing Figure(s); 9 Drawing Page(s)

LINE COUNT: 4339

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to a method for treating catabolic, qut-associated pathological processes including intestinal mucosal and pancreatic atrophy and enhanced gut permeability, impairment of host defenses and compromised immune function, and for promoting recovery from bone marrow transplantation in an animal, which comprises administering to an animal a therapeutically effective amount of glutamine or an analogue thereof.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCL INCLM: 514/563.000 NCL NCLM: 514/563.000

L15 ANSWER 10 OF 13 USPATFULL

ACCESSION NUMBER: 95:22925 USPATFULL

> 308-4994 Searcher : Shears

Use of glutamine to reduce rate of pathogenic TITLE:

microorganism infection

Smith, Robert J., Brookline, MA, United States INVENTOR(S):

Wilmore, Douglas, Brookline, MA, United States

Brigham and Women's Hospital, Boston, MA, United PATENT ASSIGNEE(S):

States (U.S. corporation)

DATE NUMBER KIND

US 5397803 US 1993-51941 19950314 PATENT INFORMATION: 19930426 (8) APPLICATION INFO.:

RELATED APPLN. INFO.: Continuation of Ser. No. US 1992-845819, filed on

9 Mar 1992, now abandoned which is a

continuation-in-part of Ser. No. US 1989-360839, filed on 2 Jun 1989, now abandoned which is a continuation-in-part of Ser. No. US 1986-906530, filed on 12 Sep 1986, now patented, Pat. No. US 4857555 which is a continuation-in-part of Ser. No. US 1985-775214, filed on 12 Sep 1985, now abandoned

Utility DOCUMENT TYPE: Granted FILE SEGMENT:

Raymond, Richard L. PRIMARY EXAMINER:

LEGAL REPRESENTATIVE: Sterne, Kessler, Goldstein & Fox

NUMBER OF CLAIMS: 1 EXEMPLARY CLAIM:

12 Drawing Figure(s); 9 Drawing Page(s) NUMBER OF DRAWINGS:

4187 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to a method for treating catabolic, AB qut-associated pathological processes including intestinal mucosal and pancreatic atrophy and enhanced gut permeability, impairment of host defenses and compromised immune function, and for promoting recovery from bone marrow transplantation in an animal, which comprises administering to an animal a therapeutically

effective amount of glutamine or an analogue thereof.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCLM: 514/563.000 INCL NCLM: 514/563.000 NCL

L15 ANSWER 11 OF 13 USPATFULL

94:47041 USPATFULL ACCESSION NUMBER:

Polynucleotides that encode the human TITLE:

proteoglycan peptide core of the effector cells

of the immune response

INVENTOR(S):

Stevens, Richard L., Sudbury, MA, United States Weis, John H., Salt Lake City, UT, United States Nicodemus, Christopher F., Franconia, NH, United

States

Brigham And Women's Hospital, Boston, MA, United PATENT ASSIGNEE(S):

States (U.S. corporation)

NUMBER KIND US 5317085 19940531 PATENT INFORMATION: US 1992-909644 19920707

APPLICATION INFO.: Division of Ser. No. US 1991-635544, filed on 18 RELATED APPLN. INFO.:

Jan 1991, now patented, Pat. No. US 5171674 which

is a continuation-in-part of Ser. No. US

1988-224035, filed on 13 Jul 1988, now abandoned

DOCUMENT TYPE: Utility Granted FILE SEGMENT:

Patterson, Jr., Charles L. PRIMARY EXAMINER: Bugaisky, Gabriele E. ASSISTANT EXAMINER:

Sterne, Kessler, Goldstein & Fox LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: 2 EXEMPLARY CLAIM: 1

8 Drawing Figure(s); 9 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT: 1550

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention relates to the identification, characterization, and sequencing of genetic sequences of human secretory granule proteoglycan peptide core protein, recombinant DNA clones directed against this sequence and against the sequence of the antisense RNA, and antibodies which recognize the native human secretory granule proteoglycan.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCLM: 530/326.000

INCLS: 530/806.000; 530/300.000

NCL NCLM: 530/326.000

NCLS: 530/300.000; 530/806.000

L15 ANSWER 12 OF 13 USPATFULL

92:102990 USPATFULL ACCESSION NUMBER:

. Polynucleotides that encode the human TITLE:

proteoglycan peptide core of the effector cells

of the immune response

Stevens, Richard L., Sudbury, MA, United States INVENTOR(S):

Weis, John H., Salt Lake City, UT, United States Nicodemus, Christopher F., Franconia, NH, United

States

Brigham and Women's Hospital, Boston, MA, United PATENT ASSIGNEE(S):

States (U.S. corporation)

NUMBER KIND DATE ______ US 5171674 19921215 PATENT INFORMATION: 19910118 (7) APPLICATION INFO.: US 1991-635544 WO 1989-US3051 19890713 19910119 PCT 371 date 19910119 PCT 102(e) date

Continuation-in-part of Ser. No. US 1988-224035,

RELATED APPLN. INFO.: filed on 13 Jul 1988, now abandoned

DOCUMENT TYPE: Utility Granted FILE SEGMENT: Wax, Robert A. PRIMARY EXAMINER: Bugaisky, Gabriele E. ASSISTANT EXAMINER:

LEGAL REPRESENTATIVE: Sterne, Kessler, Goldstein & Fox

9 NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1

8 Drawing Figure(s); 9 Drawing Page(s) NUMBER OF DRAWINGS:

1546 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention relates to the identification, characterization,

and sequencing of genetic sequences of human secretory granule proteoglycan peptide core protein, recombinant DNA clones directed against this sequence and against the sequence of the antisense RNA, and antobodies which recognize the native human secretory granule proteoglycan.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCL INCLM: 435/069.100

INCLS: 435/070.100; 435/320.100; 435/240.100; 435/240.200; 435/252.300; 435/252.330; 536/027.000; 935/066.000;

935/070.000; 935/072.000

NCL NCLM: 435/069.100

NCLS: 435/070.100; 435/252.300; 435/252.330; 435/320.100;

435/355.000

L15 ANSWER 13 OF 13 USPATFULL

ACCESSION NUMBER: 92:20916 USPATFULL

TITLE: Generation and selection of novel DNA-binding

proteins and polypeptides

INVENTOR(S): Ladner, Robert C., Ijamsville, MD, United States

Guterman, Sonia K., Belmont, MA, United States Kent, Rachel B., Wilmington, MA, United States

Ley, Arthur C., Newton, MA, United States

PATENT ASSIGNEE(S): Protein Engineering Corporation, Cambridge, MA,

United States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 5096815 19920317

APPLICATION INFO.: US 1989-293980 19890106 (7)

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Schwartz, Richard A.

ASSISTANT EXAMINER: Ulm, John D. LEGAL REPRESENTATIVE: Cooper, Iver P.

NUMBER OF CLAIMS: 42 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 12 Drawing Figure(s); 12 Drawing Page(s)

LINE COUNT: 8344

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Novel DNA-binding proteins, especially repressors of gene expression, are obtained by variegation of genes encoding known binding protein and selection for proteins binding the desired target DNA sequence. A novel selection vector is used to reduce artifacts. Heterooligimeric proteins which bind to a target DNA sequence which need not be palindromic are obtained by a variety of methods, e.g., variegation to obtain proteins binding symmetrized forms of the half-targets and heterodimerization to obtain a protein binding the entire asymmetric target.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

INCL INCLM: 435/069.100

INCLS: 435/172.300; 435/252.300; 435/320.100

NCL NCLM: 435/069.100

NCLS: 435/252.300; 435/320.100

(FILE 'HCAPLUS, MEDLINE, BIOSIS, EMBASE, WPIDS, CONFSCI, SCISEARCH, JICST-EPLUS, JAPIO, CABA, AGRICOLA, VETU, VETB, USPATFULL' ENTERED

AT 11:27:21 ON 10 APR 2003) - Author (S) 5298 S "JACOBS A"?/AU L16 458 S "VERMEIJ P"?/AU L17 2 S L16 AND L17 L18 5754 S L16 OR L17 L19 2 S L19 AND INTRACELLULARIS L20 2 S L18 OR L20 L21 1 DUP REM L21 (1 DUPLICATE REMOVED) L22 L22 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2003 ACS DUPLICATE 1 2002:503432 HCAPLUS ACCESSION NUMBER: 137:77871 DOCUMENT NUMBER: Cloning of genes for novel Lawsonia TITLE: intracellularis outer membrane proteins and their use in preparing vaccines for porcine proliferative enteropathy Jacobs, Antonius A. C.; Vermeij, INVENTOR(S): Paul Akzo Nobel N.V., Neth. PATENT ASSIGNEE(S): Eur. Pat. Appl., 26 pp. SOURCE: CODEN: EPXXDW DOCUMENT TYPE: Patent English LANGUAGE: FAMILY ACC. NUM. COUNT: 1 PATENT INFORMATION: KIND DATE APPLICATION NO. DATE PATENT NO. ____ -----_____ EP 1219711 A2 20020703 EP 2001-204919 20011214 EP 1219711 A3 20021106 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR JP 2003000276 A2 20030107 JP 2001-385373 20011219 A5 20020627 AU 2001-97371 20011220 AU 2001097371 EP 2000-204660 A 20001220 PRIORITY APPLN. INFO.: The present invention relates i.a. to nucleic acid sequences encoding novel Lawsonia intracellularis proteins. It furthermore relates to DNA fragments, recombinant DNA mols. and live recombinant carriers comprising these sequences. Also it relates to host cells comprising such nucleic acid sequences, DNA fragments, recombinant DNA mols. and live recombinant carriers. Moreover, the invention relates to proteins encoded by these nucleotide sequences. The invention also relates to vaccines for combating Lawsonia intracellularis infections and methods for the prepn. thereof. Finally the invention relates to diagnostic tests for the detection of Lawsonia intracellularis DNA, the detection of Lawsonia intracellularis antigens and of antibodies against Lawsonia intracellularis.

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